

Dechlorane Plus and its *syn*- and *anti*-isomers

Draft risk profile

Prepared by the intersessional working group of the
Persistent Organic Pollutants Review Committee

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Table of contents

[TOC \o "1-3" \u]

[PAGE * MERGEFORMAT]

Executive summary

1. At its fifteenth meeting, the Persistent Organic Pollutants Review Committee concluded that Dechlorane Plus (DP, CAS No. 13560-89-9) and its *syn*-isomer (CAS No. 135821-03-3) and *anti*-isomer (CAS No. 135821-74-8) fulfilled the screening criteria in Annex D of the Convention (decision POPRC-15/2), and decided to establish an intersessional working group to review the proposal further and to prepare a draft risk profile in accordance with Annex E to the Convention. Parties and observers were invited to submit information specified in Annex E, before 2 December 2019.
2. The “Dechlorane Plus”™ technical mixture is a commercially available polychlorinated flame retardant that has been in use since the 1960s. It is used as an additive flame retardant in electrical wire and cable coatings, plastic roofing materials, connectors in TV and computer monitors, and as a non-plasticizing flame retardant in polymeric systems, such as nylon and polypropylene plastic.
3. The technical DP mixture contains two stereoisomers, *syn*-DP and *anti*-DP, that are present in ratios of about 1:3 or 25 % *syn*-DP and 75 % *anti*-DP, and may contain low levels of a number of impurities. DP and its isomers are only known intentionally produced.
4. DP production by Hooker Chemicals and Plastics Corporation (now the Occidental Chemical Company (OxyChem)) in Niagara Falls, New York started in the 1960s. DP has been considered as a high production volume chemical in the United States with an estimated annual production of 450-4500 tonnes from 1986. Production at the Anpon Electrochemical Co. manufacturing plant in China started in 2003 and the annual production is reported to be 300–1000 tonnes per year.
5. DP is released to the environment during production, processing and use, as well as from waste disposal and recycling activities. Releases from use include industrial uses as well as releases from consumer products. DP has been detected globally in many locations, ranging from production and recycling sites to urban, rural and remote areas. DP has been detected in air, water, ice, soil, wastewater, sludge, biosolids, landfill leachate, indoor and outdoor dust, wildlife and humans.
6. On a global scale, the highest DP concentrations were detected in the United States and China close to known production sites or electronic waste (e-waste) treatment facilities. Monitoring studies document the long-range transport of DP to remote regions via the atmosphere, ocean currents and possibly also via migratory birds. Modelling studies suggest DP has transport and persistence properties similar to listed POPs. DP has been detected in different environmental matrices and biota in the Arctic, Antarctic and mountain regions of Tibet. The atmospheric half-life of DP is calculated to ~14 hours, i.e. below the criterion of two days set in Annex D (d) (iii) of the Convention. However, modelled half-lives in air are largely based on gas phase reactions and do not consider possibly longer half-lives following sorption to particles, which is presumed to be the primary mode of transport for DP due to its low vapour pressure and a high log octanol-air partitioning coefficient. Available monitoring data confirms this assumption and shows that DP is found predominantly in the particulate phase both in air and water. Long-range transport of DP is thus likely mediated by particle binding. Sorption to particles may slow down reaction rates, increase the actual half-lives in air and facilitate the long-range transport of DP.
7. DP has very low water solubility and a high octanol-water partition coefficient. It is expected to bind to organic matter in soil and sediments and is therefore probably not easily bioavailable for microorganisms. Furthermore, DP has no functional groups that are susceptible for hydrolysis and is not expected to hydrolyse. Modelling predicts that aerobic biodegradation of DP would be very slow and similar to analogue chemicals (mirex, chlordane, heptachlor, dieldrin/endrin, aldrin and endosulfane) that are already listed under the Stockholm Convention. DP is detected in deep soil and sediment cores, indicating its persistence.
8. Bioconcentration factors above 5000 have recently been reported for DP in fish studies. Depuration half-lives corresponding to a bioconcentration factor above 5000 L/kg have also been reported for DP, as well as biomagnification and trophic magnification factors above 1 in a variety of food chains and locations, including an Antarctic food chain. DP has been identified as a very persistent and very bioaccumulative substance (vPvB) in the European Union.
9. Exposure of the general population to DP takes place by consumption of food and drinking water, inhalation of indoor and ambient air, as well as respiratory and oral uptake of dust containing DP. DP has been detected in human blood and breast milk in many regions. Moreover, the human foetus may be exposed to DP via the umbilical cord blood, and breast milk may be an important source of exposure for infants. The highest DP levels have been observed in occupationally exposed workers and residents living near production facilities and e-waste recycling sites in China.
10. Available animal studies suggest low concern for acute toxicity via oral, inhalative or dermal routes of exposure in humans and other organisms, but the toxicity data on DP is limited and chronic toxicity studies are missing. Toxicity studies with DP report effects such as oxidative damage, neurotoxicity and potential for endocrine disruption. Oxidative stress has been observed in marine macroalgae, fish, marine bivalves, earthworm, birds, and

mice. In a marine macroalgae exposed to DP, photosynthesis was reduced at low concentrations. Neurotoxicity was observed in fish and potential neurotoxicity indicated in earth worms. Potential endocrine modulative effects have been described in fish and some studies indicate possible endocrine modulative effects in humans. DP has also been reported to cross the blood-brain barrier and to be maternally transferred to offspring in several species. A study in zebrafish embryos indicate synergistic effects on neurotoxicity parameters and elevated bioaccumulation of both compounds as a synergistic effect of co-exposure to DP and the 3-methyl phenanthrene.

11. DP is widely detected in the global environment, including in remote regions. Available scientific data show that DP is persistent, bioaccumulative, and potentially toxic to animals and humans. It is transported to locations far from production sites and places of use. Therefore, it is concluded that DP, with its *syn*- and *anti*-isomers, are likely, as a result of their long-range environmental transport, to lead to adverse human health and/or environmental effects such that global action is warranted.

1. Introduction

1. In May 2019, Norway submitted a proposal to include the chlorinated flame retardant (FR) Dechlorane Plus (CAS No. 13560-89-9) and its *syn*- (CAS No. 135821-03-3) and *anti*- (CAS No. 135821-74-8) isomers in Annexes A, B and/or C to the Stockholm Convention on Persistent Organic Pollutants (POPs). The proposal (UNEP/POPS/POPRC.15/3) was submitted in accordance with Article 8 of the Convention and reviewed by the Persistent Organic Pollutants Review Committee (POPRC) at its fifteenth meeting in October 2019

2. The commercial mixture “Dechlorane Plus”TM is additive FR that has been in use since the 1960s (Shen et al 2011). For Dechlorane Plus the acronyms DP or DDC-CO are commonly used. In this risk profile, DP is used.

Chemical Identity

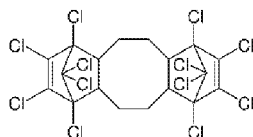
3. The technical mixture “Dechlorane Plus”TM (CAS No. 13560-89-9) is a commercially available formulation that contains two stereoisomers, *syn*-DP (CAS No. 135821-03-3) and *anti*-DP (CAS No. 135821-74-8) in a ratio of about 1:3 or 25 % *syn*-DP and 75 % *anti*-DP (Sverko et al., 2011). According to the North American manufacturer OxyChem, their commercial product contains approximately 35% *syn*-DP and 65% *anti*-DP (OxyChem, 2013). *f*_{*syn*} values for commercial DP from the Anpon Electrochemical Co and OxyChem manufacturers are in the range 0.20-0.41 (Wang et al., 2010a).

4. The chemical identity and the modelled and experimental physico-chemical properties for DP and its two isomers are listed in Tables 1 and 2, below. The structural formula of DP and its two isomers is shown in Figure 1 below.

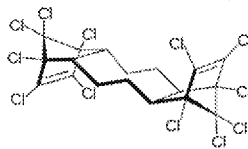
Table 1: The chemical identity of Dechlorane Plus, with its *syn*- and *anti*-isomers.

CAS number:	13560-89-9, 135821-03-3, 135821-74-8
IUPAC name:	1,6,7,8,9,14,15,16,17,17,18,18-dodecachloropentacyclo-[12.2.1.1 ^{6,9} .0 ^{2,13} .0 ^{5,10}] octadeca-7,15-diene
EC number:	236-948-9
EC name:	1,6,7,8,9,14,15,16,17,17,18,18-Dodecachloropentacyclo[12.2.1.16,9.02,13.05,10] octadeca-7,15-diene
Molecular formula:	C ₁₈ H ₁₂ Cl ₁₂
Molecular weight:	653.73 g/mol
Synonyms:	Bis(hexachlorocyclopentadieno)cyclooctane; 1,2,3,4,7,8,9,10,13,13,14,14-Dodecachloro- 1,4,4a,5,6,6a,7,10,10a,11,12,12a-dodecahydro- 1,4:7,10-dimethanodibenzo[<i>a,e</i>]cyclooctene; Dodecachlorododecahydrodimethanodibenzocyclooctene
Trade names	Dechlorane Plus 25 (Dech Plus); Dechlorane Plus 35 (Dech Plus-2); DP-515; Dechlorane 605; Dechlorane A; DP; Escapeflam DK-15 (China); PyroVex SG (grade 515, 25 and 35)

Dechlorane Plus
(CAS no. 13560-89-9)



Anti- (or *exo*) Dechlorane Plus
(CAS no. 1358 21-74-8)



Syn- (or *endo*) Dechlorane Plus
(CAS no. 135821-03-3)

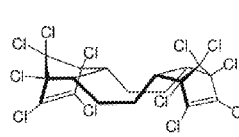


Figure 1. Structural formula of DP and its two isomers

Table 2. Overview of selected physicochemical properties of Dechlorane Plus, with its *syn*- and *anti*-DP isomers

Property	Value	Reference
Physical state at 20 °C and at 101.3 kPa	Solid white powder	ECHA, 2017b
Melting/freezing point	340-382 °C 350 °C	ECHA, 2017b OxyChem datasheet (2007)
Vapour pressure	0.006 mm Hg \pm 0.8 Pa (at 200 °C)	OxyChem datasheet (2007)
Water solubility*	<1.67 ng/L (20 – 25 °C) 0.044 – 249 μ g/L (insoluble)	ECHA, 2017b OxyChem datasheet (2007)
n-Octanol/water partition coefficient, K_{ow} (log value)	9.3	OxyChem datasheet (2007)
n-Octanol-air partition coefficient K_{oa} (log value)	12.26	OxyChem datasheet (2007)
Sediment/water partition coefficient K_p (log value)	6.65	OxyChem datasheet (2007)
Air-water partition coefficient K_{aw} (log value)	The following log K_{AW} values are estimated at 25 °C -3.2 (from measured water solubility and estimated vapour pressure) 0.44 (from measured water solubility and vapour pressure) -2.8 (from EPIWIN **predicted water solubility using log K_{OW} of 9 and vapour pressure) -3.5 (from HENRYWIN***v.3.20, predicted from structure using Bond Method).	ECHA, 2017b

*There is some uncertainty in the precise value for water solubility (Chou et al. (1979a). However, all available measurements and predictions agree that the substance is very poorly water soluble.

The following modelling programs are individual models in EPI Suite™:

** Estimation Program Interface Suite for Windows

*** Calculates the Henry's Law constant (air/water partition coefficient) using both the group contribution and the bond contribution methods. Since the group contribution method is not applicable to DP due to fragments missing in the database, only the result of the bond contribution method is shown.

5. DP is produced by the Diels-Alder reaction of two moles of hexachlorocyclopentadiene with one mole of 1,5-cyclooctadiene. The commercial technical hexachlorocyclopentadiene may contain low levels of a number of impurities such as tetrachloroethylene, pentachlorocyclopentenone, octachlorocyclopentadiene hexachlorobutadiene, and the listed POPs hexachlorobenzene, pentachlorobenzene and mirex, (US.EPA, 1978). DP monoadduct (DPMA) may be formed through partial reactions during the synthesis of DP through the diadduct Diels–Alder process and can therefore be present as an impurity in commercial DP (Sverko et al., 2010). DP compounds with lower number of chlorine atoms such as mono-dechlorinated DP (DP-1Cl or DP-Cl₁₁) may also occur as impurities in the commercial substance (Li et al. 2013b).

6. Quantitative structure-activity relationship (QSAR) modelling (Sverko et al., 2011; Feo et al., 2012) and testing done by the manufacturer (EHSI, 2004) indicate that DP has physical-chemical characteristics typical of other POPs, including high lipophilicity (log K_{ow} ~9), resistance to biodegradation, and bioaccumulation (log BAF ~5 in fish) and long-range environmental transport (LRET) (EHSI, 2004; Sverko et al., 2011; Feo et al., 2012). DP is structurally similar to substances already listed in the Stockholm Convention such as aldrin, mirex, and chlordane. Physicochemical properties of similar structures listed in Stockholm Convention, dechlorane-related substances and dechlorinated DP are described in Table 1-2 in the UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION].

Conclusion of the POPs Review Committee regarding Annex D information

7. At its fifteenth meeting, the POPs Review Committee evaluated the proposal by Norway to list DP (CAS No. 13560-89-9) and its *syn*-isomer (CAS No. 135821-03-3) and *anti*-isomer (CAS No. 135821-74-8) under the Stockholm Convention on Persistent Organic Pollutants and concluded that DP and its *syn*-isomer and *anti*-isomer met the screening criteria specified in Annex D (UNEP/POPS/POPRC.15/7), Decision POPRC-15/4). The Committee decided to review the proposal further and to prepare a draft risk profile in accordance with Annex E to the Convention.

Data sources

8. The draft risk profile is based on the following data sources:

- a) The nomination report submitted by Norway (UNEP/POPS/POPRC.15/3);
- b) Annex E information and comments submitted by Parties and Observers according to Annex E of the Convention: Belarus, Canada, Egypt, Germany, Hungary, Monaco, Netherlands, New-Zealand, Romania, Thailand, Qatar, Republic of Korea, State of Palestine, International Pollutants Elimination Network (IPEN) and Alaska Community Action on Toxics (ACAT);
- c) Reports and other grey literature as well as information from peer-reviewed scientific journals.
- d) Documents supporting the identification of DP as a Substance of Very High Concern (SVHC) under the Registration, Evaluation, Authorisation and Restriction of Chemicals (REACH) regulation in the European Union (EU) (ECHA, 2017b,c);
- e) Screening assessment report and risk management approach from Canada (Canada, 2019a,b).

Status of the chemical under international conventions

9. DP is currently not known to be included in any international conventions, however regulatory processes has been initiated in several countries.

10. DP is listed on Canada's Domestic Substances List (DSL) (ECCC, 2019). A final screening assessment of DP was published by Environment and Climate Change Canada (ECCC) and Health Canada in spring 2019 (Canada, 2019a). The assessment concludes that DP meets the criteria for toxicity to the environment, as it is entering or may enter the environment in a quantity or concentration or under conditions that have or may have an immediate or long-term harmful effect on the environment or its biological diversity. The proposed regulatory approach is to amend the *Prohibition of Certain Toxic Substances Regulations, 2012* to prohibit the manufacture, import, use, sale and offer for sale of DP and all products containing the substance (Canada, 2019b).

11. In the EU, based on an Annex XV dossier and a Risk Management Options Analysis prepared by the United Kingdom, DP (including its *syn*- and *anti*-isomers) was identified as a SVHC and added to the REACH Candidate List in 2018 as it is considered to be very persistent and very bioaccumulative (ECHA, 2017a).

12. In Norway, DP was added to the list of priority substances in January 2019 with a national goal to phase out the use by 2020 (Norwegian Environment Agency, 2019b).

13. Certain restrictions, approval and notification requirements for imports and use are also in place in United States (US), New Zealand, Thailand (see Table 3 of UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]).

2. Summary information relevant to the risk profile

2.1 Sources

2.1.1 Production, trade, stockpiles

14. DP has been manufactured by Hooker Chemicals and Plastics Corporation (now Occidental Chemical Company (OxyChem)) in Niagara Falls, New York in the US since the 1960s (Shen et al., 2011). DP has been classified as a high production volume chemical (>500 tonnes) in the US (Sverko et al., 2011), and the annual productions were estimated to be 450–4500 tonnes since 1986 (Qiu et al., 2007). Manufacturing in China by Anpon Electrochemical Co. in Huai'an has been reported to be 300–1000 tonnes per year since 2003 (Wang et al., 2010a). The global annual production is approximately 5000 tonnes (Ren et al., 2008).

15. In the EU, DP has been registered under REACH by two registrants (one for each manufacturer) with an annually tonnage band 100-1000. The major registrant since 2013, Occidental Chemical Belgium BVBA, representing the manufacturer OxyChem US, submitted a notification in December 2017 of "cease of manufacture or import" under REACH and no longer produces or imports DP to the EU (ECHA, 2020). The Dutch company ADAMA Agriculture BV has since 2016 been is registrant for the sole importer/distributor Velsicol, who represents Anpon Electrochemical Co. In 2019, imports into the EU was below 100 tonnes, and highest imported tonnage bands was 300-400 t/y (information from the Netherlands). Sweden registered use of 5 tonnes DP in 2006 and 11 tonnes in 2005 (Kaj et al., 2010).

16. In Canada, based on surveys conducted under section 71 of CEPA for the years 2011 and 2016, between 1000 and 10 000 kg of DP, including DP in some products, was imported by a few companies (Canada, 2019a,b). In the Republic of Korea, based on information gathered from government surveys (2010, 2014 and 2016); DP was not manufactured in the Republic of Korea. No DP was imported in 2010, but use was around 40 tonnes. In 2016, 11 tonnes were imported, and 17 kg was used. DP has not been exported since 2010, when 0.9 tonnes of DP was exported (Annex E information, Republic of Korea).

2.1.2 Uses

17. DP originates exclusively from anthropogenic sources. DP is currently marketed as an alternative/replacement for commercial decabromodiphenyl ether (c-decaBDE) and as a substitute for mirex used as FR (Hoh et al., 2006). It is used as a FR in electronic wiring and cables, automobiles, plastic roofing materials, and hard plastic connectors in televisions and computer monitors, wire coatings, and furniture (Zhang et al., 2015). It is also used extensively in the aerospace industry as a FR and in other applications (AIAC, 2020). DP is often used as an additive to various formulations and can be found in many aircraft components such as cabin interior panels, ducting, engines and other components. In a recent study from China, DP was detected in various building materials (Hou et al., 2018). Samples of wallpaper (non-woven, PVC, paper), latex paint, boards (e.g. laminated floor, fiber board, solid wood), glue, sealant, PVC line pipes and foam (such as sound absorbing foam and expanded polystyrene panels) contained DP in the range n.d. – 5.1 ng/g (Hou et al., 2018). In Canada, DP is used Canada as a FR in automobile manufacturing (Canada, 2019b). Annex E information from The Netherlands confirms use in automobiles in the powertrain, cooling, chassis and bodywork parts.

18. DP is used in many polymeric systems as a FR. These systems are typically either thermoplastics or thermosets. Examples of thermoplastics that may contain DP include nylon (Weil and Levchik 2009; KEMI, 2019), polyester (KEMI, 2019), acrylonitrile butadiene styrene (ABS), natural rubber, polybutylene terephthalate (PBT), polypropylene, and styrene butadiene rubber (SBR) block co-polymer (OxyChem, 2007). DP may be used in thermosets such as epoxy and polyester resins, polyurethane foam, polyethylene, ethylene propylene diene monomer rubber, polyurethane rubber, silicon rubber, and neoprene (OxyChem, 2007). The amount of DP in these materials ranges from 8% in PBT to up to 40% in silicon rubber (OxyChem, 2007). According to the manufacturer (OxyChem, 2007), DP is manufactured for use solely by industrial customers. Under REACH, DP is registered for industrial use in the following product categories; polymer preparations and compounds, semiconductors, adhesives and sealants. The following sectors of end use are listed: 1) Manufacture of plastics products, including compounding and conversion, 2) manufacture of computer, electronic and optical products, electrical equipment and 3) manufacture of plastics products, including compounding and conversion. In addition, use in glued articles in aircraft parts are indicated (ECHA, 2020).

19. In 2011, a 220-tonne sample of representatively composed e-waste was processed in a Swiss recycling facility with the different output streams being analysed. The highest DP mass flows were in fine-grained plastics and in monitor and laptop casings (Taverna et al., 2017).

20. DP was detected in plastic casings from two TVs and one computer, but not in circuit boards from similar equipment collected from a recycling facility in China (Li et al., 2019c). In addition, DP was detected in recycled ABS from Guangdong Province in China (Cao et al., 2020).
21. PyroVex® SG is an additive FR that, according to the supplier, contains > 99% DP (Velsicol Chemicals LLC, 2020). PyroVex® SG is typically mixed into plastics and other materials and can be used alone or be formulated with complementary chemistries to meet performance requirements. It is marketed for use in flame retarded wire and cable, as a FR for electrical connectors used in computers and other electronic devices. PyroVex® SG grade 515, 25 and 35 is the same chemical mixture only with different average particle size.
22. During a public EU consultation on the ninth draft recommendation for inclusion of DP in Annex XIV of REACH, held from September to 5 December 2018, a number of companies identified applications in which they use DP as an additive FR. Identified applications were; in the aviation and automotive industries, in polymeric compounds for cable insulation and minor use within the development of fireworks (ECHA, 2017e).

2.1.3. Releases and emissions to the environment

23. DP and its isomers are not known to be unintentionally produced. The occurrence of DP and its isomers in the environment is a result of human activities.
24. A number of activities such as production, use, recycling and waste handling, as well as landfill leachate and run-off, wastewater treatment), can lead to releases of DP to the environment (Wang et al., 2016; Wang and Kelly, 2017). DP has been detected in sludge (e.g. de la Torre et al., 2011; Barón et al., 2014a; Norwegian Environment Agency, 2018b; 2019a; Ji et al., 2018) and biosolids (Davis et al., 2012) from wastewater treatment plants (WWTPs). In addition, sludge from WWTPs contaminated with DP can be used as soil enrichment (Wu et al., 2017) or fertilizer (Dai, 2011 reviewed in Ji et al., 2018). Furthermore, *syn*- and *anti*-DP was detected in the particulate fraction of storm water in Oslo, Norway (Norwegian Environment Agency, 2018b; 2019a).
25. Samples from Canadian WWTPs were investigated for chlorinated FRs. Detection frequency of DP was greater than 90% in both wastewater and biosolids samples, while median removal efficiencies were between 51% and 66% for total DP (Shanmuganathan et al., 2018). In a conventional WWTP in Shanghai DP levels were detected in wastewater with a mean concentration of 0.46 ng/L (range: 0.05 to 1.40 ng/L). Sludge contained DP in the range of 1.1 to 2.0 ng/g. The fraction of *anti*-DP (f_{anti}) was consistently higher than that of *syn*-DP (f_{syn}). Average f_{syn} values ranged from 0.16 to 0.33 and was in the range of two commercial DP mixtures. The annual release of DP via sewage sludge from all WWTPs in Shanghai were estimated to be 164.8 g (Xiang et al., 2014).
26. DP is used as a FR in electrical and electronic equipment and recycling of e-waste is reported to be a source of DP release to the environment (Wang et al., 2016). Very high levels of DP in soil (3327 ng/g) was detected at a recycling site in China suggesting that e-waste recycling activities serve as an important source of DP emissions in China (Yu et al., 2010). In a Swiss study, the overall DP concentration in e-waste was 33±11 mg/kg. This translates into an annual DP mass flow of 2.3±0.9 tonnes/year in all Swiss e-waste generated in 2011 (Taverna et al., 2017).
27. Indoor dust is the sink and carrier of many pollutants including DP released from consumer products. Indoor dust is also considered to be the main source of pollutants in WWTPs receiving only domestic wastewater (Katsoyiannis and Samara, 2004; Xiang et al., 2013). DP has been detected in indoor dust in the range 0.08 to 124 000 ng/g (Newton et al., 2015; Wong et al., 2017; Cao et al., 2014; Lee et al., 2020). For additional information on releases and emissions from manufacture, WWTPs, e-waste recycling, landfills etc., see Table 4 and 13 in UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION].

2.2. Environmental fate

2.2.1 Persistence

DP is chemically stable in various environmental compartments with minimal or no anaerobic degradation (reviewed in Wang et al., 2016). DP does not contain any functional groups that are susceptible to hydrolysis and hydrolysis is not expected to be a relevant degradation process (Canada, 2019a; ECHA, 2017b). Due to a very low water solubility and high log K_{ow} , DP is not likely to be bioavailable for microorganisms and is expected to bind to organic carbon in soil and sediments (Wang et al., 2016). There are no measured half-life data for degradation of DP in surface water, sediment or soil (ECHA, 2017d). However, physical–chemical properties of DP were predicted using three different models EPISuite, SPARC and Absolv and the estimated half-lives in water, soil and sediment were predicted to 180, 3650 and 1621 days, respectively (Zhang et al., 2016).

28. Studies on photodegradation of DP in air (Sverko et al., 2008; Wang et al., 2011; Li et al., 2013b; Wang et al., 2013b; Tao et al., 2015) and water (Chou et al., 1979b) are reviewed in ECHA (2017d). Although photodegradation and formation of lower chlorinated dechloranes and other degradation products are reported in some of the studies, DP is considered to be relatively photo-stable in air under natural conditions. First, sorption of DP to

airborne particles is expected, which would lower the photolysis rate and result in a longer half-life in air (Canada, 2019b). Second, in the study of Wang et al. (2013b), where the photodegradation of DP was investigated by irradiation using a xenon lamp, rapid photodegradation by dechlorination was found under 200–750 nm light. The degradation became much slower in 280–750 nm light. The quantum yields of dechlorination by-products at 200–280 nm (UV-C) were about 2–3 orders of magnitude higher than at 280–320 nm, and no yields were detected in the 320–750 nm range (Wang et al., 2013b). As the majority of UV exposure in the environment is UV-A light with a wavelength in the 315–400 nm range, these findings suggest that DP should be photo-stable in the environment and only undergo limited degradation under natural light conditions at the terrestrial surface. However, DP degradation under UV-C irradiation was so fast that photodegradation would be a potential removal process. The sequential degradation process where one to four chlorine (Cl) are removed from DP and exchanged with hydrogen (H) suggested that the reaction was leading to photodechlorination of DP (Wang et al., 2013b). Dechlorination was also observed by Li et al. (2013b). *Anti*-DP, *syn*-DP, and commercial DP solutions were exposed to UV light. Reported dechlorination products were mainly $[-1\text{Cl}+1\text{H}]$, $[-2\text{Cl}+2\text{H}]$ (Li et al., 2013b). The only available study on photodegradation in an aqueous medium suggests that photolytic degradation in water is a possibility (Chou et al., 1979b), but the findings cannot be related to the aquatic environment because the test solution included acetonitrile and the initial concentration of DP was 1 ppm [1 mg/L], which is significantly greater than the reported solubility in pure water (ECHA 2017d).

29. Predictions using the BIOWIN v4.10 model (US EPA, 2012), Catalogic (2012) and TOPKAT (2004) suggest that biodegradation of DP will be very slow (Canada, 2019a; ECHA, 2017d). ECHA (2017d) concludes, based on the estimated data from the BIOWIN v4.10 model and associated uncertainties, that "DP is unlikely to be biodegradable". The BIOWIN results obtained for DP are the same as a number of hexachloro-norbornene-containing analogue chemicals and indicate that DP are as persistent as analogue chemicals such as mirex, chlordane, heptachlor, dieldrin/endrin, aldrin and endosulfan that are already listed as POPs under the Stockholm Convention. Furthermore, the two possible microbial degradation pathways predicted for DP are the same as for the POP analogues. According to ECHA (2017d) it is unlikely that metabolic rates for these pathways will be more rapid for DP, which is significantly less water soluble than the already listed POPs. Despite some uncertainties BIOWIN v4.10 is generally considered as a valid model for DP. The validity of the BIOWIN v4.10 model is discussed in detail in ECHA (2017d).

30. In line with these modelling results laboratory tests discussed in Canada (2019a) have shown that DP is not likely to biodegrade under aerobic conditions; an activated sludge biodegradation test (modified MITI OECD 301C) reported 0.6% biodegradation in 2 weeks (US EPA, 2011), and a 21-day test using wastewater biosolids found 0% biodegradation (US EPA, 2009).

31. Monitoring studies provide additional evidence that DP is persistent and that biodegradation in sediment is limited. The study of Qiu et al. (2007) suggests that DP can still be present over thirty years after initial deposition. In this study, DP was measured in a sediment core from central Lake Ontario, Canada, and detected in a layer corresponding to around 1980. A linear trend ($r_2 = 0.739$) of increasing f_{anti} values with time was observed, from an average of 0.76 in surficial (recent) sediments to >0.90 in the deeper layers corresponding to around 1980, suggesting that the *anti*-DP could be more persistent than the *syn*- isomer in sediment. However, the variation of f_{anti} in commercial batches over this time period is not known, and the findings could also reflect the isomerisation of *syn*-DP to *anti*-DP. Similar to the findings of Qiu et al. (2007) other studies on DP and f_{anti} values in freshwater- (Wang et al., 2010) and marine sediment (Fang et al., 2014), respectively report possible stereoselective depletion of *syn*-DP in freshwater sediment and an enrichment of *anti*-DP. The enrichment of *anti*-DP is explained as likely being due to preferential biodegradation of the *syn*- isomer in the sediment.

32. Degradation of DP in aging soils has recently been investigated. A slight decrease in DP concentrations suggested that only limited degradation occurred during the study, with only 4.2–8.2% of initial DP having degraded after 260 days (Cheng et al., 2019). Following 260 days of aging of soil containing three different concentrations of DP (0.1, 1, 10 $\mu\text{g/g}$ DP soil), 89.6% to 98.8% of the initial DP concentration was detected suggesting that only a small proportion of DP might degrade in soil over time under the influence of aging. Model simulations were then developed to investigate the transformation. The model simulations indicated that transformation rates were inconstant and statistically different from one another over time, and half-lives of DP were estimated to range from 1325 to 2948 days, further indicating its environmental persistence in aging soils (Cheng et al., 2019). It has also been reported that *syn*-DP varied little with depth down to 100 cm in soil sampled close to the Chinese manufacturing facility (Wang et al., 2010a), possibly suggesting that isomer-specific degradation or preferential adsorption does not play a significant role in soil. The DP concentration was highest in surface soil (0–5 cm; 837 $\mu\text{g/kg}$ dry weight (dw)), decreasing to 9.16 $\mu\text{g/kg}$ dw at a depth of 60–70 cm and 3.84 $\mu\text{g/kg}$ dw at a depth of 90–100 cm. The f_{anti} value varied little with soil depth but ranged from 0.75 in the surficial layer to 0.67 in the deepest layer (90–100 cm). The study authors make conflicting statements about their finding. On the one hand, they say their findings imply a stereoselective depletion of the *syn*- isomer in soil in comparison with the measured f_{anti} value of the Chinese commercial product, at 0.60. On the other hand, they also state that isomer specific microbial degradation or preferential adsorption does not play a significant role in soil.

33. Modelling studies of degradation potential and microbial metabolic pathways suggests that biodegradation of DP is likely to be very slow, and that the probability that DP will degrade any faster than analogue substances listed under the Stockholm Convention is low. Monitoring data shows limited degradation in soil and sediments over time and support the conclusion that DP is very persistent.

2.2.2 Bioaccumulation and pharmacokinetics

34. The log K_{ow} for DP is reported to be 9.3 (OxyChem, 2007). The high log K_{ow} and the very low water solubility (Table 2) indicates that DP is very hydrophobic and partitions to organic matter. This is further supported by the relatively high log sediment-water partition coefficient (log K_p) of 6.65. These properties make it difficult to perform aqueous laboratory studies as it is difficult to maintain stable exposure levels. Therefore, aqueous exposure is expected to be of limited importance in terms of bioaccumulation potential. However, significant uptake has been shown to occur in fish in a number of field studies (Guo et al., 2017; Malak et al., 2018; Kurt-Karakus et al., 2019).

35. A recent laboratory-scale microcosm fish study reported bioconcentration factor (BCF) values exceeding the bioconcentration criteria set by the Stockholm Convention (BCF >5000) (Wang et al., 2019). In this study, carp were exposed continuously for 32 days to DP (range 0.14 to 0.24 ng/L) through water, followed by 32 days depuration. Equilibrium was reached and BCF was calculated based on wet weight concentrations in muscle and ratio of uptake and elimination constants. The reported BCF values were 5700 and 9300 L/kg for *syn*- and *anti*-DP, respectively and depuration half-life were 6.3 and 7.2 days for *syn*- and *anti*-DP, respectively (Wang et al., 2019).

36. Depuration half-life is an important measure of bioaccumulation. A depuration half-life around 8-10 days is indicative of a lipid-normalised and growth-corrected BCF above 5 000 L/kg according to the analysis in Environment Agency (EA), UK (2012). In a dietary bioaccumulation laboratory test using juvenile rainbow trout, the depuration half-life (whole body minus liver) for DP was reported to be 30-40 days for the *anti*-isomer and 50-70 days for the *syn*-isomer, which is highly indicative of a very bioaccumulative substance (Tomy et al., 2008). Results from an aquatic food web study from Antarctica (Na et al., 2017) observed in line with Tomy et al., (2008) higher bioaccumulation of the *anti*-DP isomer. However, a laboratory study in carp showed tissue-specific variations (Tang et al., 2018). Furthermore, a comparison of modelled studies (Larisch and Goss, 2018) and measured (Tomy et al., 2008) oral uptake in fish showed good agreement, indicating that bioaccumulation of super hydrophobic chemicals like DP via food can be reliably predicted. Uptake of DP is slow but will eventually result in substantial bioconcentration as the substance is not metabolised (Larisch and Goss, 2018).

37. In rats exposed to commercial DP 25 by gavage for 90 days at different doses (0.1, 10, and 100 mg/kg/d), DP preferentially accumulated in liver rather than muscle. At high doses (10 and 100 mg/kg/d) *syn*-DP was predominant in tissues. The elimination half-life for *syn*-DP was about 179 days in liver, 44 days in muscle and 24 days in serum, and for *anti*-DP, 54 days in muscle and 25 days in serum. Depuration time for *anti*-DP for the liver was not reported (Li et al., 2013b).

38. Several field studies report biomagnification factors (BMF) and trophic magnification factors (TMF) for DP in various food webs. Factors, such as, temperature, time of sampling, reproduction status, migration, age and tissue versus whole body calculations may affect the calculation of TMF (Borgå et al., 2012; Franklin, 2016). DP biomagnifies in various food webs, both from freshwater and marine waters, as BMFs (Tomy et al., 2007; Wu et al., 2018; Sun et al., 2015; Sun et al., 2017) and TMFs (Sun et al., 2015; Kurt-Karakus et al., 2019; Na et al., 2017) are reported to be greater than 1, including in one study in Antarctica (Na et al., 2017). Bioaccumulation and maternal transfer have been observed in amphibians collected from a highly contaminated site in South China (Wu et al., 2018). Amphibians occupy an important trophic position in the food web between aquatic organisms and terrestrial biota. Based on the known predator-prey relationship between frog and insects, BMFs above 1 (1.8 – 2.7) for DP were reported in this study (Wu et al., 2018). In a study of biomagnification of DP in a freshwater reservoir in the vicinity of an electronic recycling facility in South China, the TMFs of the DP isomers were found to be 2-3 times greater than those of PBDE congeners and comparable to or lower than those of the highly recalcitrant PCB congeners in the same food web (Wu et al. 2010). In an aquatic food web study from China, TMFs were reported in the range 1.9 – 3.1. In this study, biomagnification potential (TMF >1) were also reported for DPMA, *anti*-Cl₁₁-DP and *anti*-Cl₁₀-DP (Wang et al., 2015), and this should also be taken into account when evaluating bioaccumulation for DP. For more information on these substances, see section 1.1 Chemical identity. Three terrestrial studies are available showing BMFs above 1 (Yu et al., 2013; She et al., 2013; Wu et al., 2018). For further details on the bioaccumulation studies, see Table 5 in UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION].

39. The isomer composition of DP in environmental samples can be different from the technical products because of their biota isomer-selective uptake or elimination, bioaccumulation, biodegradation, and stereospecific photodegradation (Wang et al., 2015; and reviewed in Wang et al., 2016). Trophic levels (Peng et al., 2014; Tomy et al., 2007; Wang et al., 2015), exposure concentrations (Li et al., 2013a, b; Mo et al., 2013; Peng et al., 2015), life cycle stages (Klosterhaus et al., 2012; Sührling et al., 2014), and the type of wildlife and tissue (Peng et al., 2014; Zhang et al., 2011; Zheng et al., 2014a) and stereo-selective excretion (Li et al., 2019) are the main factors for the stereoselective bioaccumulation of DP in organisms.

40. Studies in bird indicate that uptake, distribution and elimination kinetics of DP and its isomers are complex. Chickens bred in the vicinity of an e-waste recycling facility exposed to DP through the environment from sand and food showed preferential accumulation of the *anti*-DP isomer (Zheng et al., 2014a). Lipid contents of the tissue was the main factors in the tissue distribution, although degree of blood perfusion state and tissue function impacted too. Furthermore, the study revealed tissue specific accumulation of *anti*-DP, with higher *anti*-DP levels in brain, fat and liver (0.065-0.064) compared to (0.54-0.59) for other tissues. The study also showed that *anti*-Cl₁₁-DP was absorbed through the diet rather than metabolised from DP *in vivo* (Zhen et al., 2014a). This information is supported by the fact that *syn*-Cl₁₁-DP and *anti*-Cl₁₁-DP were detected in commercial DP-25, hence, it can be inferred that these chemicals originated from commercial products (Li et al., 2013b).

41. There are some studies on biotransformation of DP (Hoh et al., 2006; Tomy et al., 2007; 2008; Zhu et al., 2014). DP has been shown not to metabolise easily in biota (Tomy et al., 2008; Xian et al., 2011) and the results from Tomy et al. suggest enzyme-induced metabolism of DP in biota might be low, if it does occur. Degradation products of DP such as decachloropentacyclooctadiene (DP-Cl₁₀) or undecachloropentacyclooctadiene (DP-Cl₁₁) have been detected in bird eggs (Guerra et al., 2011; Muñoz-Arnanz et al., 2011, 2012; Zheng et al., 2014a), but some studies suggest they are formed through biotic or abiotic processes prior to uptake or even through impurities in the commercial product (Sverko et al., 2008, 2010; Tomy et al., 2008; Zheng et al., 2010, 2014b). Two additional dechlorination analogues was also detected in both quails and the test substance (Li et al., 2013a,b). In a study where eggs from Japanese quail were injected in the yolk sac with DP, no biotransformation of DP was shown (Briels et al., 2018).

42. DP was first detected in archived fish (walleye) from Lake Erie sampled from 1980-2000, which suggested that DP was taken up by the fish (Hoh et al., 2006). Available monitoring studies show that DP is widely dispersed in the environment (reviewed in Canada, 2019a; ECHA 2017b). Global evidence reveals that uptake of DP can occur in various biota samples such as aquatic and terrestrial food webs as well as the human body (reviewed in Wang et al., 2016). Field monitoring data suggest that DP is bioavailable and can exceed levels in biota that are of concern based on critical body burden considerations related to baseline narcosis (ECHA 2017b). DP was detected in human milk samples collected in two Canadian cities (Siddique et al., 2012), in human serum (Brasseur et al., 2014; Yan et al., 2012; Ren et al., 2009; Chen et al., 2015) and cord serum (Ben et al., 2014) as well as in human hair (Chen et al., 2015). Furthermore, very high DP concentrations in blood and hair from workers of a manufacturing facility have been reported (Zhang et al., 2013), see Table 10 and 13 in UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]).

43. In summary, a long depuration half-life, which is indicative of a BCF above 5000 L/kg, has been reported for DP. BCF above 5000 L/kg have been reported for DP in fish studies as well as BMFs and TMFs > 1 for several organisms and food webs, and this further supports the conclusion that DP is bioaccumulative. Many studies have revealed the presence of DP in wildlife and humans. Field monitoring data suggests that DP is bioavailable and can achieve high body burdens (ECHA, 2017b). Additionally, DP was recently identified as a very bioaccumulative (vB) substance in the EU (ECHA, 2017a).

2.2.3 Potential for long-range transport

44. Due its high hydrophobicity and low vapour pressure (see Section 1.1), DP in the atmosphere adsorbs to airborne particles (Sverko et al., 2011; AMAP, 2017, Canada, 2019a). Monitoring studies have reported mean fractions of particle-bound DP in air as high as 99 % (Hoh et al., 2006). Measurements in seawater confirm the propensity of DP to adsorb to particles. In the East-Greenland Sea, particle-bound DP accounted for, on average, 97% of *syn*-DP and 80% of *anti*-DP. In Atlantic seawater, the particle bound fraction was on average 58% for *syn*-DP and 75% for *anti*-DP (Möller et al., 2010).

45. The modelled half-life (QSAR estimate) of DP in air is 13.68 hours (Sverko et al., 2011), i.e. below the criterion of two days set in Annex D (d) (iii) of the Convention. However, half-lives in air are largely derived using models that are based on gas phase reactions and do not take into account possibly longer half-lives following sorption to particles, which is presumed to be the primary mode of transport for DP (reviewed in Sverko et al., 2011; Canada, 2019a). As shown already for other chemicals with similar physico-chemical properties such as decaBDE (Brevik et al., 2006; UNEP/POPS/POPRC.10/10/Add.2.; POPRC-10/2), sorption to particles may slow down reaction rates, increase the actual half-life in air and facilitate LRET of DP on particles (Sverko et al., 2011; Canada, 2019a). As described in ECHA (2017d), the LRET of substances with low vapour pressure that adsorb strongly to particulates in the air is likely to be governed by the fate of the particulates to which they bind and can undergo LRET to remote regions when atmospheric conditions permit (e.g. during dry periods).

46. Modelling studies performed with the OECD Pov and LRTP Screening Tool, a software tool for screening chemicals for persistence and LRET potential (Wegmann et al., 2009), and using α -HCH, HCB, PCB-101, PCB-180, PCB-28, HBCD, atrazine, *p*-cresol, chlordene plus, Dec 602, Dec 603 Dec 604 as benchmark chemicals, suggest DP has transport and persistence properties within the range for listed POPs (Sverko et al., 2011), and that DP may be deposited to some degree in remote regions (Canada, 2019a). However, the results generated using the OECD

Pov and LRTP Screening Tool are associated with uncertainty, largely because most of the input parameters used in the model are estimated (ECHA, 2017d; Sverko et al., 2011).

47. DP has been detected in many different environmental matrices and biota in remote regions; in Arctic air, snow, soil, sediment, water and biota (reviewed in AMAP, 2017; Vorkamp et al., 2019a,b; Canada, 2019a); in Antarctic air, soil, lichens, mosses, penguins, skuas, southern elephant seals and Antarctic fur seals (Möller et al., 2010; Gao et al., 2018; Kim et al., 2015; 2018; Aznar-Aleman et al., 2019); and in air, soil and lichen in a remote mountain region in Tibet (Yang et al., 2016a; Liu et al., 2018).

48. In Arctic biota, most detections have been made in species in the marine food web (polar bear, ringed seal, harp seal, hooded seal, beluga whale, killer whale, narwhal, glaucous gull, common gull, black guillemot eggs, common eider, European shag, kittiwake, mink, and cod (*anti*-DP only)). Detections also include terrestrial and freshwater species, i.e. reindeer dung, peregrine falcon eggs and landlocked Arctic char (reviewed in AMAP 2017; Canada, 2019a; see also Norwegian Environment Agency, 2018a; Dreyer et al., 2019; Letcher et al., 2018; Green 2019; Verrault et al., 2018; Vorkamp et al., 2015; 2018; 2019a; Houde et al., 2017; Simond et al., 2017; Schlabach et al., 2011).

49. As discussed in further detail in section 2.3.1 below, while concentrations in remote regions are generally low, they are not always lower than DP levels in source regions (see Table 7. UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]). Reported detection frequencies vary from non-detect to 100% detection. Collectively, the available monitoring data suggests that DP can be transported over large geographical distances to remote areas where it deposits, transfers to the receiving environment and is taken up in biota (reviewed in Sverko et al., 2011; AMAP 2017 and Canada 2019a; see also Möller et al., 2010; 2011; 2012; Na et al., 2015; Yang et al., 2013; Wang et al., 2010b).

50. Möller et al. (2010) were the first to publish environmental monitoring data demonstrating the potential for DP to undergo LRET. In this study, marine boundary layer air and surface seawater samples were collected during a sea expedition from the East-Greenland Sea via the Northern and Southern Atlantic toward Antarctica. DP concentrations ranged from 0.05 to 4.2 pg/m³ in the atmosphere and from non-detect to 1.3 pg/L in seawater. The results indicate that transport may occur both via air and seawater and show preferential partitioning to particles in both air and seawater (Möller et al. (2010)). In the East-Greenland Sea, the authors observed decreasing concentrations of DP in air with increasing latitude. The authors suggest that the finding reflect stereoselective depletion of *anti*-DP likely caused by UV degradation during LRET. The trend of declining levels with increasing latitude was not reflected in the seawater samples. In the study, Western Europe was identified as a source region for DP in the marine environment. The analysis of air mass back trajectories showed a mix of oceanic, Arctic and continental air masses. The higher levels of DP in seawater observed at some northern sites were probably due to water masses originating from the Atlantic and Arctic Oceans, in combination with freshwater inputs from melting land ice.

51. On a sea expedition from the East China Sea to the Arctic (Möller et al., 2011), DP concentrations ranged from 0.01 to 1.4 pg/m³ in air and from 0.006 to 0.4 pg/L in seawater. The Asian continent was identified as a source of DP to this marine environment, and the air-seawater exchange indicated deposition of DP from air to water. DP concentrations in air in the Chukchi/Bering Sea region were generally low compared to the source region. Decreasing DP concentrations with increasing latitude were also observed in this study; however, concentrations were near detection limits.

52. In yet another study of DP in marine boundary layer air by the same authors, atmospheric samples were collected during a sampling expedition from the East Indian Archipelago toward the Indian Ocean and further to the Southern Ocean (Möller et al., 2012). *Syn*- and *anti*-DP were detected in all marine atmospheric samples at concentrations from 0.26 to 11 pg/m³. Measured concentrations were in the range of previously reported DP concentrations in the marine atmosphere, with the highest concentrations detected in the East Indian Archipelago (see Table 6 and 7, UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]).

53. As discussed by Vorkamp et al. (2019a), the highest DP levels in Arctic air to date were detected on Greenland in 2012. Reported DP concentrations in Greenland air in 2012, 2014 and the period 2014-2016 were 6.7, 0.64 and 4.2 pg/m³, respectively (Vorkamp et al., 2015; 2019a). DP was also positively detected in atmospheric samples from other Arctic sites; Alert in the Canadian Arctic (Xiao et al., 2012), and Longyearbyen in the Norwegian Arctic (Salamova et al., 2014), and Little Fox Lake in the Canadian Sub-Arctic (Yu et al. 2015). Mean DP concentrations reported for these sites were ~0.75, ~1.2 and ~0.25 pg/m³, respectively. DP was also detected in air from Pallas in Northern-Finland at a mean concentration of 0.039 pg/m³ (Haglund et al., 2016).

54. LRET of DP via air is also documented by studies from other remote regions than the Arctic. The studies by Möller et al. (2010;2012) demonstrate ongoing long-range atmospheric transport of DP to the Antarctic and further show that the DP levels in Antarctic air are comparable to those of other FRs such as polybrominated diphenyls (PBDEs) and polybrominated biphenyls (PBBs). These findings are supported by detections of DP in Antarctic biota as further described in section 2.3.1 Environmental levels and trends (see Gao et al., 2018; Kim et al., 2015; 2018; Aznar-Aleman et al., 2019). DP also undergoes long-range atmospheric transport to remote mountain areas. In a

study on lichen from the southeast Tibetan Plateau, DP concentrations appeared to decrease with increasing altitude (Yang et al., 2013). According to the authors, the distribution pattern for DP observed in this study, combined with knowledge about the monthly average surface wind vector field, provides evidence that DP transport into the area is mainly driven by the Indian monsoon and can be attributed to LRET. Yang et al. (2013) further note that the lower DP levels observed at higher altitudes indicate that DP transported into the area via the atmosphere is cold trapped by the mountains. DP has been detected in muscle, guano, eggs, plasma, preen oil and feathers in different species of birds around the world (Gauthier and Letcher 2009; Guerra et al., 2011; Munoz-Arnanz et al., 2011; 2012; Baron et al., 2014b; 2015; Vorkamp et al., 2018; Løseth et al., 2019; Briels et al., 2019; Mo et al., 2019; Desjardins et al., 2019).

55. Birds have previously been identified as biovectors for the transport and deposition of POPs to ecosystems in remote regions through deposition of guano, feather loss and decaying carcasses (Evenset et al., 2007) and may represent an additional transport pathway for DP to remote regions. In Ellasjøen at Svalbard, seabird guano was found to account for approximately 14% of the POPs contaminant inventory of the lake catchment area, approximately 80% of the POPs contaminant inventory of the lake itself and was found to be approximately thirty times more efficient as a contaminant transport pathway for POPs compared to atmospheric LRET (Evenset et al., 2007).

56. In a study by Desjardins et al. (2019), deposition of guano by urban-adapted ring-billed gulls from a colony on Deslauriers Island near Montreal, Canada were postulated to increase exposure locally (Desjardins et al., 2019). The total amount of halogenated FRs deposited by the entire ring-billed gull colony (64 980 gulls, both sexes combined) through guano was estimated to 1 g during a 28-day period.

57. Vorkamp et al. (2018) detected DP in peregrine falcon eggs collected in Kujalleq in South-Greenland in the low Arctic. Greenland peregrine falcons are migratory birds that spend the summer in the northern hemisphere and the winter in the Caribbean and South America. Thus, the DP in their eggs could reflect exposure experienced at wintering and breeding grounds, as well as during migration.

58. As discussed, Möller et al. (2010) postulated that the fraction of *anti*-DP in environmental samples (i.e. $f_{anti} = anti/(syn + anti)$ based on concentrations) decreases with increasing distance from the source as a result of more rapid degradation of the *anti*-DP isomer than the *syn*-DP isomer in UV-light. However, these findings may also reflect isomerisation of the *anti*- to *syn*-DP (Sverko et al., 2011). Available monitoring studies provide some evidence of changes in isomer ratios with increasing distance from the source (ECHA, 2017b), but observations from the Arctic are generally inconclusive in this regard (AMAP, 2017). More specifically, DP isomer ratios in Arctic air have been reported in some studies to be in the same range as found in commercial mixtures (Xiao et al., 2012; Salamova et al., 2014; Vorkamp et al., 2015), but have in other studies (Möller et al., 2010; 2011; Carlsson et al., 2018; Na et al., 2015), been reported to differ from the commercial mixtures. Findings from Antarctica are consistent with a change in DP isomer ratio during LRET. In a study from the Fildes Peninsula in Antarctica, Gao et al. (2018) estimated the mean f_{anti} values in soil and lichen to 0.37 and 0.24, respectively. These f_{anti} values are lower than those of commercial products ($f_{anti} = 0.64-0.80$) and the authors conclude that the findings confirms that long-range atmospheric transport is a main source of DP, and that the DP burdens could be driven by a stereoselective enrichment of *syn*-DP during atmospheric transport as postulated by Möller et al. (2010). However, as the concentration of DP in soil and lichen differed between the sampling sites, the authors also suggest that the DP levels may have been influenced by anthropogenic and animal activities in the area (Gao et al., 2018).

2.3 Exposure

2.3.1 Environmental levels and trends

59. DP is detected worldwide in many different environmental matrices and biota, and at different types of locations spanning from production sites and recycling facilities to urban, rural and remote areas (Sverko et al., 2011; Wang et al., 2016). The highest DP concentrations were generally detected in the US and China with concentrations up to several ng m⁻³, close to known production or e-waste recycling sites (Sverko et al., 2011).

60. DP levels in remote regions and in the oceans are generally much lower than levels reported in source regions near production sites and urban centres, although high levels in some instances also have been observed at remote sites (reviewed in Sverko et al., 2011; Canada, 2019a). As further detailed below, due to limited data, the temporal and spatial trends for DP are equivocal.

61. Several DP monitoring studies have been carried out in the vicinity of DP manufacturers and e-waste recycling sites in the US and in China (Sverko et al., 2011; Ji et al., 2018). High DP levels have been reported in various matrices from these areas (Wang et al., 2016). The soil concentration of DP near the Anpon Electrochemical Co. production facility in China was 1200 ng/g and the amount decreased by an order of magnitude within 7.5 km (Wang et al., 2010c). A very high DP concentration (3327 ng/g) was found at an e-waste recycling site in Qingyuan, China, while DP levels fell dramatically with increasing distance away from the recycling site (Yu et al., 2010). In the US, DP has been detected in Niagara River suspended sediment samples, near the OxyChem manufacturing facility. Examination of sediments collected between 1980 and 2006 showed a declining concentration trend for DP ($R=0.60$, $p<0.01$) (Shen et al., 2011). A declining concentration trend for DP was also reported previously for Niagara River

suspended sediments (Sverko et al. 2008). and are consistent with declines observed in a sediment core collected near the river mouth on the Niagara River bar (Sverko et al., 2010). The suspended sediment trends observed by Shen et al. (2011) were also reflected in open lake sediment cores, which generally showed a decline in DP concentrations since 1980 in the Niagara basin. Similar findings were reported for the Mississauga basin (Shen et al., 2010). It is not clear if the decline in DP in Niagara River suspended sediment samples represent changes in production volumes at the manufacturing site or whether it reflects emission controls adopted over the past 20 years at the site.

62. In a study on halogenated FRs in eels from five Latvian lakes, nine dechlorane-related compounds including mirex, *syn*-DP, *anti*-DP, Dec 602, Dec 603, Dec 604, hexachlorocyclopentadienyldibromocyclooctane, Cl₁₀DP and Cl₁₁DP were analyzed. The mean total concentration of dechlorane-related compounds was 0.62 ng/g lw and the geographical distribution was nearly uniform among the five lakes (Zacs et al. 2018). Dec 602 was the predominant accounting for approximately 40% of the total concentration of dechlorane-related compounds. The concentrations of *syn*- and *anti*-DP ranged from non-detectable to 0.45 ng/g lw, and from non-detectable to 0.89 ng/g lw, respectively, with average concentrations of 0.14 ng/g lw, and 0.24 ng/g lw, respectively. The contribution of *anti*-DP to the total concentration of dechlorane-related compounds was approximately 33%, while *syn*-DP accounted for approximately 10% on average. The composition of mixtures containing *syn*- and *anti*-DP showed an f_{anti} value that was close to the composition of the OxyChem® DP commercial product, indicating the release of DP from disposal of consumer products as the probable source. Urban-feeding ring-billed gulls breeding in a highly industrialized section of the St. Lawrence River, downstream from Montreal (QC, Canada) was found to contain high levels of DP. The DP hepatic mean level was 230 ng/g lw. *Anti*- and *syn*-DP were detected in 100% and 93% of the investigated livers, respectively (Gentes et al., 2012).

63. As described above and in section 2.2.3, DP is detected in remote areas globally. See also Tables 6-9 in UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]. A review of emerging Arctic contaminants conducted by AMAP concluded that atmospheric concentrations of DP in the Arctic were comparable to those of PBDEs (AMAP, 2017). While DP concentrations in remote regions are generally low, they are not always lower than in source regions. For example, in the study investigating halogenated FRs in air and soil from Mt. Gongga on the eastern Tibetan plateau, DP levels ranging from below detection limit to 11.5 pg/m³ and 8.3 pg/g in air and soil, respectively, were reported (Liu et al., 2018). The Tibetan Plateau is the world's highest elevation plateau and is considered as one of the most remote and isolated areas on earth, and ideal for the study of LRET of POPs (Yang et al., 2016a). According to the authors (Liu et al., 2018), the concentrations of DP in air and soil on the Tibetan plateau were higher than in the Great Lakes region (0.14–4.0 pg/m³) (Sverko et al., 2011), but within the range of air samples from China (not detected to 66 pg/m³) (Ren et al., 2008), and much lower than those observed near a production facility in China (7740–26,700 pg/m³) (Wang et al., 2010a). In another study on lichen from the southeast Tibetan Plateau, DP was detected in 89% of the samples (Yang et al., 2016a). Concentrations ranged from 20 to 1121 pg/g, with an average concentration of 318 pg/g. The concentrations reported were comparable, but slightly higher than in tree bark from Shenzhen, Hangzhou and Tianjin in China (Qiu and Hites, 2008) and much lower than in tree barks from South Korea and New York in the US (Qiu and Hites, 2008) but significantly higher than in moss from Ny-Ålesund at Svalbard (Na et al., 2015).

64. In Antarctic soil, lichen and moss samples, DP was detected in all samples (100%) (Gao et al., 2018; Kim et al., 2018). The DP concentrations in Antarctic soil were between one and several orders of magnitude lower than the levels reported in surface soil samples collected from an industrial region (0.0336–4.65 ng/g) and an e-waste recycling area (nd–47.4 ng/g) in South China (Yu et al., 2010). On the other hand, DP levels in both lichen and moss from Antarctica were higher than previously measured in mosses sampled in Ny-Ålesund in Arctic Norway in 2012 (Na et al., 2015). The average concentrations of DP in lichen from Antarctica were similar to those in reeds in northeastern China (0.63 ng/g dry weight (dw)) (Wang et al., 2012) and in lichen in the southeast Tibetan Plateau (167 pg/g) (Yang et al., 2016a) but lower than in tree bark from areas in the northeastern US (0.03–115 ng/g) and South Korea (1.4 ng/g) (Qiu and Hites, 2008).

65. In a Norwegian screening assessment of emerging Arctic contaminants, DP was detected in all species monitored. The detection frequency within the species tested was 100%, except in common eiders where it was 80% (Norwegian Environment Agency, 2018a). *Syn*- and *anti*-DP concentrations in organisms from Svalbard were in the range from non-detectable to 6.9 ng/g ww with reported average values of 0.04 and 0.07 ng/g ww for common eider (eggs), 0.01 and 0.03 ng/g ww for kittiwakes (eggs), 0.24 and 1.8 ng/g ww for glaucous gull (eggs), 0.63 and 4.5 ng/g ww for polar bear (blood). Moreover, DP was also detected in European shag (eggs) from Røst, in mink (liver) from Sommarøy and in common gulls from Tromsø. Reported *syn*- and *anti*-DP levels in these species were 0.22 and 1.4 ng/g ww in European shag, 0.21 and 1.2 ng/g ww in mink, and 0.1 and 0.63 ng/g ww in common gulls, respectively. In another study conducted for the Norwegian Environment Agency, DP levels were below the limit of quantification in cod livers from Arctic sites at Svalbard and Kjøfjorden in the Outer Varangerfjord and the Tromsø harbour area (Green et al., 2019). In contrast, reported *syn*- and *anti*-DP in cod livers from the Bergen harbour area, a non-Arctic site on the west coast of Norway, were 0.178 and 0.203 µg/kg ww. *Syn*- and *anti*-DP levels in cod livers from the Inner-Oslofjord in Southern Norway were 0.135 and 0.231 µg/kg ww, respectively.

66. In a report by Vorkamp et al. (2019) on DP in air and biota from Greenland, *anti*-DP isomer was detected in 92% of all air samples from 2016 and in 46% of the air samples collected in 2014. Except for glaucous gull, DP levels

were close to or below limits of quantification in biota samples. Neither of the two DP isomers were found in ringed seal samples from Thule. In ringed seals from East-Greenland, Arctic char and narwhals, only the *anti*-DP isomer was present. *Syn*- and *anti*-DP were both detected in 100% of all glaucous gull samples from Thule at mean concentrations of 0.22 and 0.89 ng/g lw (0.012 and 0.049 ng/g ww), respectively. Detection frequencies for *syn*- and *anti*-DP in glaucous gull samples from East-Greenland were 87.5 and 100%, with mean concentrations of 0.24 and 0.93 ng/g lw for *syn*- and *anti*-DP. A geographical comparison based on samples of glaucous gull and ringed seal showed relatively uniform DP levels. The DP levels measured in this study were generally comparable to concentrations reported for some of the same species (e.g. ringed seals, glaucous gull and Arctic char) from other locations in the Arctic. However, DP concentrations in glaucous gull liver samples from Greenland reported in this study were a factor of 5-10 lower than in corresponding samples from the Canadian Arctic (Verreault et al., 2018). In the study from the Canadian Arctic, *syn*- and *anti*-DP were detected in 65% and 59% of male liver samples from 2012, respectively (Verreault et al., 2018). In females, the detection frequency of both isomers was 21%. Mean concentrations were only calculated for male birds and were 0.18 and 0.16 ng/g ww for *syn*- and *anti*-DP, respectively.

67. DP was also previously detected in air and biota samples from Greenland along with other FRs (Vorkamp et al., 2015). Mean *syn*- and *anti*-DP concentrations in air were 2.3 and 5.2 pg/m³. While the detection frequency of both isomers in air was 46%, *anti*- and *syn*-DP was detected in 95% and 100% of biota samples, respectively. In biota, ringed seals from East-Greenland had the highest DP concentrations with reported mean blubber concentrations of 0.096 and 0.42 ng/g ww for *syn*- and *anti*-DP, respectively, but were not statistically different from West-Greenland ringed seal samples. Reported mean concentrations for *syn*- and *anti*-DP in other species were; 0.14 and 0.67 ng/g ww in black guillemot eggs, 0.023 and 0.11 ng/g ww in glaucous gull liver, 0.019 and 0.071 ng/g ww in ringed seal blubber from West-Greenland and 0.021 and 0.055 ng/g ww in polar bear adipose tissue.

68. In another study, covering multiple locations in the Canadian Arctic, DP was only detected sporadically in ringed seals (Houde et al., 2017). Detectable levels of *syn*- and *anti*-DP ranged between 0.04 and 0.41 ng/g lw and 0.04-6.3 ng/g lw, respectively.

69. Minke whales and beluga whales from the St. Lawrence Estuary as well as beluga from the Canadian Arctic have also been studied (Simond et al., 2017). In 2013, the mean DP concentration for these three populations was 0.31 (minke whales, St. Lawrence Estuary), 0.44 (beluga, St. Lawrence Estuary) and 1.28 ng/g lw (beluga, Arctic). The highest concentration of 1.28 ng/g lw in the Arctic beluga population was different from the generally observed contaminant distribution, which usually showed higher concentrations at more southern locations, and could not be explained. Concentrations in beluga from St. Lawrence estuary and the Canadian Arctic were roughly the same order of magnitude as observed in two killer whales from Greenland with quantifiable levels of *anti*-DP (Vorkamp et al., 2019).

70. Research on four avian species on King George Island showed that DP was present in Gentoo penguin, Adelie penguin, South Polar skua and Brown skua (Kim et al., 2015). Reported concentrations of DP were 0.250–0.329 ng/g lipid in the penguin tissues and 2.12–11.1 ng/g-lipid in the skua tissues. Dec 603 and 604 were not detected, whereas Dec 602 was detected in the highest concentrations in all samples, followed by *anti*-DP and *syn*-DP. In another Antarctic study, Gao et al. (2018), reported mean DP concentrations of 233 ± 89 , 201 ± 86 and 150 ± 147 pg/g dw in soil from three different locations (i.e. a coastal area, an inland area and Ardely Island) on the Fildes Peninsula. Reported levels in lichen from the same locations were 449 ± 213 , 337 ± 172 and 1513 ± 1376 pg/g dw, respectively. DP was detected in all samples. Kim et al. (2018) reported the presence of DP in lichen and mosses sampled from 16 different sampling sites spread across the South-Shetland Islands, King George Island and Anvers Island in Antarctica. Similar to Gao et al. (2018) and based on geographical differences in DP levels and f_{anti} values, the authors propose that LRET, human activities, melting glacier water and biological activities, e.g. from penguins, are possible sources of DP in the area (Kim et al., 2018). Aznar-Aleman et al. (2019) investigated DP and other dechloranes in dead seals from the South Shetland Islands on the Antarctic Peninsula. Only Dec 602 and *anti*-DP were found. DP was detected in adipose tissue from southern elephant seals and in Antarctic fur seals at average concentrations of 0.60 ng/g lw and 0.16 ± 0.16 ng/g lw, respectively, but were below the limit of quantification in muscle, brain and fur. Dec 602 appeared in greater concentrations than *anti*-DP, especially in adipose tissue. Reported average Dec 602 concentrations were 1.48 ± 0.35 ng/g lw (range n.d.–1.72 ng/g lw) in fat of southern elephant seal and 1.92 ± 1.66 ng/g lw (range n.d.–3.19 ng/g lw) in fat of Antarctic fur seal. In other tissues of both species, Dec 602 levels were always below 0.55 ng/g lw.

71. Several studies have detected 1,3- or 1,5-DPMA in environmental samples such as sediment and fish (e.g. Sverko et al., 2010; Tomy et al., 2013; Sühring et al., 2014; Wang et al., 2015; Wolschke et al., 2015; Rjabova et al., 2016). In addition, it is possible that failure to use a non-destructive clean-up procedure during sample preparation could lead to under-reporting of this substance (Rjabova et al., 2016). In some cases, the concentrations of DPMA-isomers were greater than the total DP concentration in the same samples (See Table 11 in UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]). 1,3-DPMA (but not 1,5-DPMA) was detected in lake trout at an average concentration of 34 ± 43 µg/kg lw, which was around ten-fold higher than concentrations of DP in the same samples reported by Tomy et al. (2007). In peregrine falcon eggs harvested in Spain and Canada, DPMA was detected in 24 out of 25 eggs, with concentrations ranging from 1.7 to 469 ng/g lw and 1.2 to 1660 ng/g

lw in peregrine falcon eggs from Spain and Canada, respectively (Guerra et al., 2011). The corresponding DP levels was in the range 0.3 to 3.6 and 7.5 to 209 ng/g lw, respectively. DPMA has also been detected in tissues (except blood) of brown skua from King George Island, Antarctic indicating potential LRET. The detection frequencies were only 21%, 11%, and 7.1% for DPMA, Dec-604 and *anti*-DP with the highest levels of 136, 165, and 8.9 pg/g dw, respectively (Wolschke et al., 2015).

72. Spatial trends of DP in remote regions were reported in five separate studies from the Arctic (Möller et al., 2010; 2011; Vorkamp et al., 2015; 2019a,b). As discussed in section 2.4.4 above, Möller et al. (2010) and (2011) observed decreasing concentrations of DP in air, but not in water, with increasing latitude. Data from Greenland show either fairly uniform levels or no spatial trends; Vorkamp et al. (2015) found that DP levels were not statistically different in ringed seal samples from East- and West-Greenland, while the Vorkamp et al. (2019a) found a fairly uniform occurrence of DP in glaucous gull and ringed seal samples from different locations.

Temporal trend data for DP are equivocal. Air samples (gas and particle phase) from Harbin, China indicated significant and increasing trends for both *syn*- and *anti*-DP in the period 2008 to 2013 (Li et al., 2016). The time it took for *syn*-DP ($t_2 = 2.3 \pm 0.37$, $r = 0.39$, $p = 0.001$) and *anti*-DP ($t_2 = 1.8 \pm 0.25$, $r = 0.43$, $p = 0.001$) concentrations to double was significantly different, but the doubling times of both DP isomers were shorter than reported for 2-ethylhexyl 2,3,4,5-tetrabromobenzoate and bis (2-ethylhexyl) tetrabromophthalate, suggesting a more rapid increase for DP than the other FRs in the study. Air parcel backward trajectories and source contribution suggested a strong input from local sources (Li et al., 2016). In a study from North America, DP concentrations in air (gas and particle phase) increased from 2005 to 2013, with doubling times of 4–6 years at three sites (Chicago, Sleeping Bear Dunes and Eagle Harbor) in the US Great Lakes area. At the two sites closest to a known production site for DP, Cleveland and Sturgeon Point, levels were stable and unchanged in the same time period (Liu et al., 2016). In a study from Canadian sites in the Great Lake Basin (Burnt Island, Egbert and Point Petre), the levels of *syn*- and *anti*-DP in air decreased from 2005 to 2013. Reported half-lives were 3.4 years at Burnt Island and 14 years at Point Petre (Shunthirasingham et al., 2018). The declines leveled off after 2010. Research involving two of the same sites included in the Liu et al. (2016) study, Olukunle et al. (2018) found decreasing *syn*-DP levels at Point Petre and increasing levels at Burnt Island. The reported half-life at Point Petre was 6.6 while the doubling time at Burnt Island was 7.6 years. The increasing levels of DP observed at Burnt Island by Olukunle et al. (2018) and in Chicago, Sleeping Bear Dunes and Eagle Harbor by Liu et al. (2016) are consistent with the results presented by Salamova and Hites (2011) who reported an increasing trend for the *syn*-DP with a doubling time of 9.5 years at US Great Lakes sites. However, although the investigated sites are in the same region, they are not identical.

73. When compared to the global data, levels in air in Chicago and at Sturgeon Point are at the higher end of the range and are similar to the average DP level (3.5 pg/m³) from Taihu, China (Qiu et al., 2010). The increasing levels of DP observed in the Great Lakes region which are reported in some studies (Li et al., 2016; Liu et al., 2016; Olukunle et al., 2018; Salamova and Hites, 2011), and in a typical urban city in China (Li et al., 2016) may indicate that DP is being increasingly used. Using a multiple linear regression model of DP concentrations to isolate the variabilities due to sampling date and population near the sampling site Olukunle et al. (2018), showed that DP concentrations in precipitation, gas and particle phase air samples from the Great Lakes were changing as a function of sampling date. This finding indicates that the input of DP to the environment in this area is continuing, presumably because its use and production are not regulated (Olukunle et al., 2018). On the other hand, and as described earlier, decreases of DP were observed in suspended sediments, a sediment core, and lake trout samples of Lake Ontario, following peaks in the 1980s (Shen et al., 2011). The reasons for the different time trends in air, precipitation and sediment in the Great Lakes area are not clear, but as indicated by Vorkamp et al. (2018) it is possible that the matrices reflect different emission sources.

74. There is also some information on time trends in biota. A recent study by Vorkamp et al. (2018), the only study available on temporal trends of DP in the Arctic, reported DP concentrations in the range 0.984 to 37.9 ng/g lw during the period 1986 to 2014 in peregrine falcon eggs from South Greenland. A non-significant increasing trend (annual change 3.3%) was observed (Vorkamp et al., 2018). In belugas from the St. Lawrence Estuary, DP increased from 1997 to about 2000 and decreased subsequently, possibly with a second peak around 2010 (Simond et al., 2017). No significant change in DP concentrations was found in eggs of white storks or black kites collected in 1999, 2003, 2011 and 2013 from the Doñana Natural Space in Spain (Báron et al., 2015). In a study investigating FRs in Indo-Pacific humpback dolphins and finless porpoises from the South China Sea, Zhu et al. (2014) found a positive temporal trend in the ratio of DP to BDE-209 between 2003 and 2011 in dolphin samples and between 2003 and 2012 in porpoise samples. The finding was by the authors attributed to an increasing use of PBDE alternatives in the South China region following the restriction of the production and use of PBDE commercial mixtures. A study reporting levels and trends of different FRs in the German and polar environments from 2005 to 2015/2016 found declining trends for DP in some of the investigated biological matrices (i.e. tree leaves, tree shoots, herring gull eggs) and increasing trends in bream file and blue mussel soft body (Dreyer et al., 2019). In the same period, a small declining trend was also observed in riverine suspended matter. However, standard deviation in this sample set was quite large. As noticed by the authors herring gulls feed in the marine environment, e.g. on blue mussels, but may also have a terrestrial diet. Thus, one possibility is that the seemingly contrasting time trends reported by Dreyer et al. (2019) reflect differences in DP emission and deposition between terrestrial and aquatic environments.

75. Rauert et al. (2018) report air monitoring data for from 48 global sites across all five UN regions that were collected using polyurethane foam passive air samplers. The study generally showed low detection for all FRs at background sites. PBDEs showed higher mean concentrations and detection frequencies than novel FRs (Rauert et al., 2018). *Syn*-DP was below the detection limit at all 48 sites and *anti*-DP at all sites except Paris, France, where high levels of DP were reported (19 in quarters one and four, 7.5 pg/m³ in quarter two and 116 pg/m³ in quarter three in 2014). When compared to monitoring data from 2005 and 2005 and 2008 to 2009, respectively, Paris had elevated concentrations of novel FRs and reduced PBDE levels. *Anti*-DP and the FRs β -tetrabromoethylcyclohexane and bis(2-ethyl-1-hexyl) tetrabromophthalate were the dominating novel FRs. Rauert et al. (2018) propose that "the increasing concentrations of the replacement FRs and the reduced levels of the PBDEs may be indicating the shift away from the use of PBDEs to alternative FRs in this urban area".

2.3.2 Human exposure

76. Humans may be exposed to DP from indoor dust, food, indoor and ambient air, water, soil, sediment and breast milk. DP has been detected in domestic dust samples from several countries indicating exposure from products and manufactured items containing DP (Shoeib et al., 2012; Johnson et al., 2013; Wong et al., 2017; See Table 12 in UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]). Higher levels of DP in air from urban compared to rural environments have been observed (Ren et al., 2008; Chen et al., 2011). A regular distribution pattern of DP in dust with particle size was observed with stronger enrichment on fine particles than coarse particles, which indicates potential human exposure (Cao et al., 2014). As reported in section 2.1.3 above, in indoor dust in Beijing, China, highest DP levels were found in pooled samples from classrooms (231 ng/g), which was higher than in house dust from Guanzhou, China (mean 18.9 ng/g) (Cao et al., 2014). In indoor dust from different microenvironments (i.e. home, workplace, car) in Greater Cairo, Egypt, *syn*- and *anti*- DP were detected in 100% of car dust samples reaching 5 and 6 ng/g, respectively (Hassan and Shoeib, 2015). In the Republic of Korea, DP was detected in almost all samples of indoor dust from homes (0.30 – 530 ng/g; median 9.1 ng/g), offices (3.7 – 100 ng/g; median 35 ng/g), and daycare centers (6.2 – 56 ng/g; median 12 ng/g) (Lee et al., 2020). Air sampling in 15 homes in India found DP levels ranging from 0.2 to 5.43 pg/m³ (median 2.81 pg/m³) and 0.52–62.7 pg/m³ (median 1.62 pg/m³) for urban and suburban sites, respectively (Yadav et al., 2020). Cequier et al. (2015) found no correlation between DP levels in indoor air, dust or serum levels for 46 Norwegian women. However, both levels in air, dust and serum were low in these samples. Mean DP levels in serum were 2.6 ng/g lw and dust samples from the corresponding homes had mean *syn*- and *anti*-DP levels of 9 and 19 ng/g, respectively while the air concentrations were even lower (Cequier et al., 2014). High levels of DP were detected in ventilation dust from aircraft cabins produced between 1986 and 2008 with a range of 31 to 9600 ng/g for *anti*-DP (Allen et al., 2013).

77. In food products from Belgium, DP was found in 5% of the 1289 samples (firm cheese, quail eggs and pork, mean DP levels were 339, 637 and 331 pg/g ww, respectively) (Poma et al., 2018). In 175 samples of 35 different food products from the retail market in the Republic of Korea, the mean DP was in the range from ND to 170 pg/g ww, and *syn*- and *anti*-DP was detected in 83.4 and 79.4% of the foodstuff, respectively (Kim et al., 2014). Daily intake of DP was estimated to be 11.2×10^3 pg/day, which was 3 orders of magnitude higher than for other dechloranes (Dec 602, 603, 604 and Mirex). The highest level of DP was found in bovine liver DP 170 pg/g ww followed by spanish mackerel, oysters and dried anchovy with 135, 81 and 78 pg/g ww, respectively. However, in this study, DP was detected in all foodstuffs except onion. This result shows the wide-spread exposure of DP in the environment of Korea (Kim et al., 2014). In a study from Lebanon of 58 food samples representing fatty food groups, the estimated daily dietary intake for the adult population had lower and upper boundaries of 1344 and 1718 pg/day for DP (Malak et al., 2019). In a market basket study of 123 food samples from Japan, DP was detected in Groups III (sugar and confectionary), V (legumes and their products e.g. soy bean, tofu, fried bean curd, natto, chickpea), X (fish, shellfish and their products), and XI (meat and eggs) at concentrations of 3.3, 2.8, 1.9, and 1.5 pg /g ww, respectively (Kakimoto et al., 2014). The atmospheric concentration in Osaka city was 11 pg/m³ and the estimated daily intake of DP (750 pg/day) via inhalation and diet was approximately one percent of that for Σ PBDE (62 ng/ day) (Kakimoto et al., 2014).

78. In a study comparing human exposure at a production facility and an e-waste recycling facility in Southern China, high levels of DP were found in foodstuffs produced in proximity to the e-waste recycling site and production facility, where vegetables contained from 305 ng/g ww (paikchoi cabbage) to 2720 ng/g ww (green onion) DP (Wang et al., 2013a). Grains (common wheat, maize and soybeans) had levels of 1370, 760 and 498 ng/g ww, respectively, and common carp and other local fish species had levels from 56.8 to 1110 ng/g ww (Wang et al., 2013a). Estimated dietary intake dose accounted for more than 99% and 93% of the daily intake to people included in the study from the production site and e-waste recycling site, respectively, which assumed that consumed food was produced locally (Wang et al., 2013a). For the production site, the estimated dietary exposure dose was 1.1×10^{-2} mg/kg/d for workers, residents and children and the estimated dietary exposure dose was lower for the e-waste recycling site at 3.0×10^{-5} mg/kg/d (Wang et al., 2013a). Total estimated exposure dose (dietary, dermal, inhalation) in people from the e-waste recycling site was 0.03×10^{-3} mg/kg/d, which is 300 times lower than at the manufacturing site (Wang et al., 2013a). In another study from an e-waste recycling area in China, free-range chicken eggs harvested in 2010, 2013 and 2016

contained high levels of FRs, including DP. No temporal trend was observed for DP levels. The highest levels of DP were observed in eggs from 2016, mean and SD was 5413 ± 1978 ng/g lw (Huang et al., 2018).

79. DP has been detected in human serum from several countries. In a study from France mean $\Sigma 5$ PBDEs (BDE-47, -99, -100, -153 and -154) levels (4.32 ± 2.99 ng/g lw) were in the range typical of Western Europe levels, but lower than the mean $\Sigma 5$ dechlorane compounds (Dechlorane 603 > DP > Mirex > Dechlorane 602 > Chlordene Plus) levels (6.24 ± 4.16 ng/g lw) (Brasseur et al., 2014). Levels of DP observed were lower in serum from Canada, France, Korea and Norway compared to occupationally exposed people in China with median of 2.39, 1.20, 0.73, 1.3 and 42.6 ng/g lw, respectively (Zhou et al., 2014; Brasseur et al., 2014; Kim et al., 2016; Cequier et al., 2015; Ren et al., 2009). In a study from China analyzing pooled human serum samples from different age groups, including 305 subjects from the south coast of Laizhou Bay, Shandong Province, China, there was no relationship between DP levels and age. However, the younger volunteers, age 20 to 29 years, had the highest serum concentrations with mean DP around 7 ng/g lw compared to mean DP for all groups 3.6 ng/g lw (Wang et al., 2014). A time trend of Dechlorane compounds was investigated in seven pooled serum samples from Korean adults to cover the period from 2006 to 2013. No distinct trend was observed for Dechlorane compounds, while PBDEs steadily decreased with time (Kim et al., 2016).

80. High levels of DP have been observed from occupational exposure at a DP manufacturing plant compared to nearby areas in eastern China. The DP concentrations ranged from 89.8 to 2958 ng/g lw in whole blood and 4.08 to 2159 ng/g dw in hair. For the workers directly engaged in the DP manufacturing process, their DP levels were significantly higher than in most individuals of the two control groups (Zhang et al., 2013). In another study, DP serum levels in 70 occupationally exposed workers from an e-waste recycling workshops and control group of 3 residents of an urban area in South China were measured. The DP levels were significantly higher in the workers (22–2200 ng/g with median of 150 ng/g lw) than in the urban residents (2.7–91 ng/g with median of 4.6 ng/g lw). The DP concentrations in females were found to be associated with their age but this relationship was not found for males (Yan et al., 2012). A study of 15 occupationally exposed workers at e-waste recycling sites in Bangladesh found a median DP exposure of 2.3 ng/h/silicone wristband, approximately 130 times higher than non-occupationally exposed US residents (Wang et al., 2020b).

81. In humans, DP has been detected in cord serum (Ben et al., 2014) and in human milk (Siddique et al., 2012) showing transfer to offspring at different developmental stages. Maternal transfer of bioaccumulative substances *in utero* represents a potential risk to embryonic development and may represent the largest source of FRs input to offspring during the first few years of life. In 450 human milk samples from Norway, the Netherlands and Slovakia, DP had detection frequencies in the range of 3 to 9% and 20 to 26% for *syn*- and *anti*-DP, respectively. The concentrations of *syn*-DP ranged between 0.05 and 2.85 ng/g lw and of *anti*-DP between 0.004 and 1.60 ng/g lw (Čechová et al., 2017). Higher detection frequency was observed for 87 human milk samples from Canada (76 and 91% for *syn*- and *anti*-DP, respectively), with mean *syn*- and *anti*-DP concentrations of 0.27 and 0.7 ng/g lw (Siddique et al., 2012). In another Canadian study of 105 milk and 102 maternal serum samples, *syn*- and *anti*-DP detection frequencies were 40 and 50% for milk samples and 77 and 78% for maternal serum samples, respectively (Zhou et al., 2014). However, concentrations of DP in milk samples were lower than in the previous study from Canada (i.e. Siddique et al., 2012). Total DP in maternal serum was 2.37 ng/g lw, which was lower than previously found in two Chinese studies (Ben et al., 2013; Ren et al., 2009), where the median value of total DP in maternal serum samples of mothers who were not exposed to e-waste recycling activities was 13.7 ng/g lw in one study (Ren et al., 2009), and 4.0 ng/g lw in the other study (Ben et al., 2013). The total DP levels in milk samples were also higher in the Chinese study with a median of 2.19 ng/g lw (Ben et al., 2013). *Syn*-DP and *anti*-DP were detected in all milk ($n = 44$) and serum samples ($n = 45$) (Ben et al., 2013). The Cl₁₁-DP detection frequency in breast milk and serum was 45% and 84%, respectively, and DP levels in blood and milk from residents who had been living in Taizhou for >20 years (R(20) group) were significantly higher than those who had living in that city for <3 years (R(3) group) (Ben et al., 2013). For overview see Table 13 in UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION].

82. In another publication from the same study of 72 residents of the e-waste recycling area of Wenling, China, DP was detected in placenta and cord serum indicating prenatal exposure of DP across the placenta (Ben et al., 2014). In contrast to human milk samples which have been reported to have a f_{anti} ratio close to that of commercial DP mixtures (0.6 - 0.8) (Zhou et al., 2014; Siddique et al., 2012; Ben et al., 2013), the concentration ratio between the cord serum and maternal serum was estimated to be 0.45 for *syn*- and 0.35 for *anti*-DP, suggesting the placenta partially limited DP transfer with a greater extent for *anti*-DP (Ben et al., 2014).

2.4 Hazard assessment for endpoint of concern

83. Available studies show that DP has the potential to elicit toxic effects in different organisms. Oxidative stress has been observed after exposure to DP in fish (Chen et al., 2017; Hang et al., 2013; Kang et al., 2016), marine bivalves (Barón et al., 2016; Gagne et al., 2017), earthworm (Zhang et al., 2014; Yang et al., 2016b), birds (Li et al., 2013a) and mice (Wu et al., 2012). The organism may be able to handle oxidative stress/damage by activating/upregulating antioxidant defense and repair mechanisms. However, reactive oxygen species are important signaling

molecules that could trigger unwanted effects by activating signaling cascades regulating cell death and cell survival and by causing damage to DNA, protein, telomeres, cells, tissues etc. Oxidative stress is linked to aging and disease and could be the extra stressor that leads to "overload" in the wildlife with multiple stressors. As further described below, reported effects include oxidative stress to most tested organisms, neurotoxicity in zebrafish and carp (Chen et al., 2017; Li et al., 2019b) and potential for endocrine modulating effects in zebrafish (Kang et al., 2016) and humans (Ben et al., 2014; Guo et al. 2019) and immune modulating effects in carp (Li et al., 2019b).

2.4.1 Toxicity to aquatic organisms

84. DP has very low water solubility (< 44 ng/L, see Table 2) and will partition to particles, sediment and biota in the aquatic environment. Observed levels in natural water may be higher than in pure rinsed water, e.g. in Laizhou Bay water, median DP was 11.7 ng/L for sum filtered and particulate, range of DP in filtered and particulate water samples were nd to 10 ng/L and 0.890 to 346 ng/L, respectively (Zhen et al., 2018).

85. Toxicity data on sediment dwelling organisms are lacking, and the Canadian screening assessment used toxicity data on the DP analogue chlordane to fill the data-gap as a worst-case-scenario for sediment dwelling organisms and estimated a predicted no effect level (PNEC) of 0.0129 mg/kg dw for DP (Canada, 2019a).

86. Two studies on aquatic algae indicate potentially negative effects on photosynthetic endpoints at low doses of DP (Gong et al., 2013; 2018). The freshwater algae *P. subcapitata* was exposed to nominal DP concentrations of 13.51, 135.1 and 1 351 ng/L for up to 72 h (Gong et al., 2013). Esterase activity had a significant but transient increase at 2 h by the two highest doses and 135.1 ng/L at 24 h. Reactive oxygen species (ROS) levels was significantly increased at all doses at 48 h, but at 72 h ROS levels in exposed cells was similar to control cells (Gong et al., 2013). At the highest dose (135.1 and 1 351 ng/L), average chlorophyll a content was significantly increased at 48 and 72 h (Gong et al., 2013). In juvenile marine green macroalgae *U. pertusa*, the uptake/ of DP affected physiological responses of photosynthesis and caused oxidative stress (Gong et al., 2018). Following a 21-day exposure (DP doses of 10^{-8} to 10^{-6} mol/L equal to ~6.53 to 653 µg/L) and 21-day depuration, *anti*-DP was prone to bioaccumulate. ΣDP accumulated (after exposure to 10^{-8} mol/L) were 127 and 206 ng/g ww on day 7 and 21, respectively. Antioxidant enzyme activities (superoxide dismutase (SOD) and catalase), oxidative damage to lipids (malondialdehyde, MDA) and chlorophyll fluorescence parameters were affected in a dose and time dependent manner after 1, 7 and 14 days of DP exposure. The effect on photosynthesis and SOD were significant at all concentrations tested, while effects on catalase and MDA were significant for the two highest doses. The authors concluded that DP affects photosynthesis in the marine algae, as observed by a low rate of light energy utilization, and that this effect may be ascribed to oxidative damage caused by the uptake of DP (Gong et al., 2018). Marine macroalgae constitute the main source of biomass production and thus support the structure and function of the coastal marine ecosystem. Adverse effects on macroalgae may also have a potential impact on other trophic levels.

87. Mediterranean mussels were exposed to DP concentrations of 0, 5.6, 56, and 100 µg/L for 6 days by spiked agarose. Concentrations measured in water immediately after dosing were 0, 0.4 ± 0.3 , 0.3 ± 0.2 and 0.7 ± 0.5 µg/L, and measured levels in mussel after 23 h exposure were 4.7 ± 3.1 , 8.8 ± 2.1 and 21 ± 9.1 µg/mussel, respectively (Barón et al., 2016). DNA strand-breaks were observed in hemocytes at all doses. No clear dose-response was observed, although all doses were significantly different from the negative control; 13, 23 and 18%, respectively. Increased micronuclei formation was observed only for highest dose (100 µg/L) tested (Barón et al., 2016). In a more long-term study, fasting blue mussels were exposed to nominal concentrations of 0, 0.001, 0.01, 0.1 and 1.0 µg DP/L for 29 days, which resulted in ΣDP tissue concentrations of 0, 0.12, 0.98, 7.26 and 57.8 ng/g ww, respectively. Following DP exposure, no histopathological lesion was found in gonads, and no change in hemocyte DNA strand breakage, phagocytosis rate, and viability was observed. Gills were identified as the most responsive tissue. Lipid peroxidation in gills was found to increase by 82% and 67% compared to control at the 0.01 and 1.0 µg DP/L dose, respectively (only significant for 0.01 µg/L), while cyclooxygenase activity (COX) was significantly decreased by 44% at the 1 µg/L dose (Gagné et al., 2017). The effect on COX could have an impact on the control of spawning process and thus reproduction in blue mussels (Matsutani and Nomura, 1987). These two studies indicate that DP exposure cause oxidative stress and add to the multiple stressors that mussels have to use energy on to compensate.

88. Four studies with embryo/larval zebrafish with waterborn DP exposure have been identified (Hang et al., 2013; Noyes et al., 2015; Chen et al., 2017; Kang et al., 2016). No significant adverse effects on hatchability, survival or malformation were seen in the Noyes et al. (2015), Chen et al. (2017) and Kang et al. (2016) studies following exposure to DP at a concentration of up to 4.18 mg/L. Kang et al. (2016) exposed zebrafish embryos to nominal DP concentrations of 0.4, 0.8 or 1.6 mg/L from 4 to 144 hours post fertilization (hpf). Measured water concentrations of ΣDP were 140, 248, 267 µg/L DP at 0 h, and 28.8, 70.5 and 71.7 µg/L at 48 h, respectively. In contrast, Hang et al. (2013), observed significant time- and dose-dependent increases in malformations (spine side curve, cardiac edema, tail deformation) in zebrafish embryos exposed to DP at nominal concentrations of 0.037 to 0.37 mg/L from 8 hpf up to 7 days (168 hpf).

89. Short-term exposure studies with embryo/larval zebrafish suggest that DP can induce oxidative stress and neurobehavioral changes (Hang et al., 2013; Noyes et al., 2015; Chen et al., 2017). Noyes et al. (2015) observed

hyperactive response to dark-stimuli-activation at 120 hpf after exposure to nominal concentration of 6.4 μ M DP (4.18 mg/L). Chen et al., (2017) observed significant increase in spontaneous movements at 24 hpf for all nominal doses (15, 30 and 60 μ g/L), dose- and time-reduced distances of swimming after touch-induced stimuli, significant decreased free-swimming activity and reduced swimming speed during each dark- and light period. These neurobehavioral changes may be linked to axonal and muscular lesions. DP significantly inhibited primary motor neuron axonal growth and induced cell apoptosis and lesions in muscle fibres of the tail region of larvae in a dose dependent manner at all doses at 96 hpf. Axonal growth-related gene-expression (α 1-tubulin and gap43) was significantly increased in the highest dose (60 μ g/L), oxidative stress markers as ROS and MDA as well as apoptotic messenger ribonucleic acid (mRNA) markers were increased at the two highest doses (30 μ g/L and 60 μ g/L) (Chen et al., 2017).

90. Co-exposure of zebrafish embryos to nominal concentrations of DP (60 μ g/L) and/or 3-methyl phenanthren (3-MP) (5 or 20 μ g/L) at 6 to 96 hours post fertilization (hpf) resulted in elevated bioaccumulation of both compounds and synergistic effects on neurobehavioral abnormalities, axonal growth reduction, apoptotic markers in muscle and brain Ca^{2+} homeostasis (Chen et al., 2019). Relative ROS level in whole fish was significantly elevated compared to control from all exposure at 96 hpf, but no significant alteration was observed for co-exposure. As in the previous study (Chen et al., 2017), significant increase in spontaneous movement occurred at 24 hpf with 60 μ g/L DP, but co-exposure with 5 or 20 μ g/L 3-MP gave a significant reduction compared to control. DP and 3-MP co-exposure showed a synergistic or additive effect in reduction of swimming distances after touch-induced stimuli and free-swimming activity. Furthermore, synergistic or additive effect were observed for reduction in ventral axonal length growth, increased number of apoptotic cells in the tail region and induction of axon related gene expression. Brain Ca^{2+} homeostasis was investigated at 96 hpf, and a significant synergistic increase was observed in brain intracellular Ca^{2+} levels, and a synergistic reduction in Ca^{2+} ATPase activity was observed after co-exposure. Furthermore, a synergistic effect was observed in reduced expression of genes related to Ca^{2+} homeostasis. Relative to DP exposure alone, co-exposure with low and high-dose 3-MP increased the accumulation of DP by 14 and 82%, respectively. The larval body burden of DP (60 μ g/L) was 583.2 ± 33.5 ng/g ww while co-exposure with 5 and 20 μ g/L 3-MP increased DP levels to 665 ± 33.5 and 1061 ± 85.7 ng/g ww, respectively. Relative to 3-MP alone, co-exposure with DP (60 μ g/L) increased 3-MP levels by 45 and 47% for the 5 and 20 μ g/L 3-MP groups, respectively (Chen et al., 2019). Elevated bioaccumulation resulting from mixture exposure may represent a significant contribution of the synergistic effects observed in combined chemical exposure.

91. Studies also show that DP can cross the blood-brain barrier in fish (Zhang et al., 2011) as well as in frogs (Li et al., 2014), and *anti*-DP had a high persistence in the brain compared to the liver (Zhang et al., 2011). Stress responses and histopathology changes of the brain and liver were observed in juvenile common carp (n= 30 per group) exposed to nominal concentrations of 0, 30, 60 or 120 μ g/L DP for 1, 15 or 30 days (Li et al., 2019b). Tissue morphology (cell abnormality rate) was significantly changed in the groups exposed for 60 and 120 μ g/L for 15 and 30 days. In liver, unclear contours, vacuolisation and nuclear lysis were observed, and in the brain, abnormalities were observed in the granular layers, nuclear cell structures, disordered arrangement, microthrombic red blood cells, increased numbers of glia cells and nodulations. Liver SOD, glutathione (GSH) and MDA activity levels decreased with increasing dose and exposure time while the opposite pattern was observed in the brain (Li et al., 2019b). DP also altered expression of immune regulating genes in the brain and liver. CYP1B1 gene expression was significantly upregulated at all timepoints and concentrations in both the liver and brain. In addition, CYP2B and CYP3A1 and the apoptosis related factors bax and bcl-2 were altered in liver. These results indicate that DP exposure perturbs metabolism in the liver and brain, inhibits antioxidant enzyme activity, increases lipidperoxidation, promotes inflammation and induces cell apoptosis in juvenile carp (Li et al., 2019b). In another experiment DP-exposed carp embryos showed DNA damage, increased mortality rate, delayed hatching time, reduced hatching rate, decreased body length, and increased morphological deformities (Li et al. 2020).

92. In the study of juvenile carp describes above, both the pro-inflammatory cytokines IL-6 and IL-1 β and the anti-inflammatory cytokine IL-10 were significantly increased compared to control at all doses at 30 days in liver, while in the brain, IL-6 was significantly upregulated with the two highest concentrations, IL-1 β with highest concentration only, and IL-10 with 30 μ g/L (Li et al., 2019b). This is the first study indicating potential immune modulating effects of DP. Interesting, low dose (10 μ g/kg bw for 7 days) Dec-602 has been shown to alter immune response by dysregulate T-helper-1 and -2 balance in mice and alter immune and transmitter-related metabolism (Feng et al., 2016; Tao et al., 2018).

93. In juvenile Chinese sturgeon treated with DP at doses of 1, 10, and 100 mg/kg ww for 14 days via a single intra peritoneal dose, liver proteomics indicate that DP had effects on the generalized stress response, small G-protein signal cascades, Ca^{2+} signalling pathway and metabolic process, and induced apoptosis in the liver (Liang et al., 2014).

94. DP exposure induces oxidative stress in adult zebrafish tissues (Kang et al., 2016; Hang et al., 2013). In zebrafish orally exposed to spiked feed with DP dosed 0.25, 2.5 and 7.5 mg/g bw per day for 7, 14 and 28 days, the highest dose induced apoptosis in intestines, and a significant dose-dependent increase of SOD activity was observed

for all doses from day 1 to 28 (Hang et al., 2013). Measured concentrations in fish tissue at day 7 were 618.4, 762.8 and 1823.4 ng/g, respectively. The proteomic profile in the liver and brain was significantly altered and identified proteins were related to DNA damage, protein synthesis, immune response, cell apoptosis and cytoskeleton (Hang et al., 2013). In zebrafish receiving doses of 0, 0.3, 1, or 3 µg/g ww of DP by gavage on day 0 and 2, measured tissue concentrations were 5, 30, 44 and 420 ng/g ww, respectively, on day 6. An increase of catalase activity was observed in the liver, indicating oxidative stress response to the liver on day 6 (Kang et al., 2016). In addition, some indication of endocrine disruption potential of DP was indicated. Transcriptional responses of both thyroid and sex hormone related genes in the brain were altered, suggesting possible thyroid and sex hormone disrupting potentials of DP (Kang et al., 2016). On day 6, significant induction of CYP19b (brain type of aromatase) was observed at all doses at body residual levels that were environmentally relevant. In addition, the brain estrogen receptor, alpha mRNA, was significant elevated at the lowest dose (0.3 µg/g) only. Plasma thyroxine (T4) concentrations increased (but changes not statistically significant) along with up-regulation of corticotropin releasing hormone and TSH-β genes in brain at the highest dose (3 µg/g ww (Kang et al., 2016), which are both involved in the fine-tuning of the thyroid hormone-pathways in non-mammalian vertebrates (De Groef et al., 2006).

95. Maternal uptake and transfer of DP has been seen in several species of fish (e.g. Wu et al., 2013; Sührling et al., 2015; Peng et al., 2012; Zhang et al., 2011; Zeng et al., 2014) and in frogs (Wu et al., 2018). DP and other FRs were found in developing embryos of female sharks, demonstrating maternal transfer *in utero* (Marler et al., 2018).

2.4.2 Toxicity in terrestrial organisms

96. Oxidative stress responses and neurotoxicity have been observed in earthworms (Zhang et al., 2014; Yang et al., 2016b). No acute toxicity or significant change in body weight were observed in earthworms exposed to DP at nominal doses of 0.1, 0.5, 6.25 and 12.5 mg/kg for 28 days (Yang et al., 2016) or up to 50 mg/kg for 14 days (Zhang et al., 2014). However, oxidative stress was indicated by alteration in markers such as SOD, MDA, glutathione, glutathione-peroxidase, catalase and 8-hydroxy-2-deoxyguanosine (8OHdG) in tissue as well as tail DNA in comet assays of isolated coelomocytes. Furthermore, acetylcholinesterase (AChE) and cellulase activity of earthworms was significantly reduced even by the low dose indicating potential neurotoxic effects in earthworms (Yang et al., 2016). The overall 28-d no-effect concentration (NOEC) for these biomarker responses is <0.1 mg/kg (ECHA, 2017b).

97. No acute toxicity has been observed for DP in birds, but studies indicate that DP is bioavailable and transferred to eggs. In paired samples of eggs and plasma from bald eagles from Canada, DP was more abundant in eggs compared to plasma with geometric mean of 0.28 and 0.43 ng/g ww in eggs from inland and Great Lakes, respectively, versus plasma levels of 0.02 ng/g ww (Guo et al., 2018).

98. Crump et al. (2011) studied concentration-dependent effects of DP using *in vivo* and *in ovo* toxicity approaches in domestic chicken embryonic hepatocytes and chicken embryos. DP was injected into eggs prior to incubation and no overt toxic effects were observed up to the maximum dose of 3 µM in hepatocytes, and up to the highest nominal dose of 500 ng/g/egg for pipping success of the young.

99. Li et al. (2013a) studied the effects of DP on male common quails continuously exposed to commercial DP by gavage for 90 days at dose concentrations ranging from 1 to 100 mg/kg bw/d. Mortality, body and liver weight was not altered in any of the exposure groups. Liver enzyme activity and oxidative stress were measured. The authors reported DP effects on some measures of enzyme activity (e.g., significant decrease of 7-pentoxoresorufin-O-demethylase (PROD) in all exposed groups relative to the control, significant increase in alkoxyresorufin O-dealkylase (ERND) and the antioxidant enzyme catalase in the high-dose exposed groups relative to control). Furthermore, the study found DP was more prone to accumulate in liver (vs. serum, muscle), and *syn*-DP accumulated (vs. *anti*-DP) in the two high-exposure groups.

100. The Canadian screening assessment used toxicity data from Mirex as a worst-case scenario to fill the data-gap for terrestrial (plant) organisms, with estimated DP predicted no effect concentrations (PNEC) of 0.075 mg/kg dw respectively (Canada, 2019a).

2.4.3 Human toxicity

101. Available assessments and laboratory studies with mammals suggests that, DP is not carcinogenic, mutagenic or toxic to reproduction (as reviewed in ECHA, 2017b,c; Canada 2019a). Other biomarkers of exposure and/or potential adverse effects in mammals have been reported such as oxidative stress, liver impairment and endocrine effects as described below.

102. In terms of *in vitro* genotoxicity, results from Ames assays conducted using *Salmonella typhimurium* strains (TA98, TA100, TA1535, TA 1537 and TA 1538) were negative in the presence or absence of metabolic activation (S9) (Mortelmans and Tanaka 1980, as described in OxyChem 2004b). Result from an *in vitro* mouse lymphoma assay was also negative in the presence and absence of S9 (Jotz and Mitchel 1980, as described in OxyChem 2004b). One *in vivo* genotoxicity study was identified. Mice were orally administered 0, 500, 2000 or 5000 mg/kg-bw/day of

DP via gavage for 10 days (Wu et al. 2012). Liver samples were collected for a comet assay and the genotoxicity result was negative.

103. Acute toxicity studies in experimental animals suggest low concern for acute toxicity via the oral, inhalation and dermal routes of exposure. No adverse health effects were observed in any of the identified repeated-dose oral toxicity studies, testing dose levels up to 5000 mg/kg-bw/day for 10, 28 days (as summarized in ECHA, 2017b,c; Canada 2019; OxyChem 2004b and further described below). However, there are some data gaps, for example, there are no long-term studies exceeding 90 days, which might be important given the apparently slow uptake and long elimination half-life of the substance. The dosing vehicles might also limit exposure (e.g. due to the presence of undissolved micro-crystals), such that the high doses might not truly reflect the degree of exposure of the organisms (ECHA, 2017b, c). The 5000 mg/kg bw dose was described as particularly viscous (Brock et al., 2010), and is an extremely high dose (5 times the limit dose of the OECD test guideline). Further toxicity testing (*in vitro* AMES test and a pre-natal developmental (PNDT) study has been required by ECHA following a compliance check on the REACH registration dossier. (deadline 21/12/2020).

104. Wu et al. (2012) report liver impairments in mice at high-dose exposure. Following a 10-day oral exposure, oxidative stress and damage was induced in male mouse livers at all doses (500, 2000 or 5000 mg/kg-bw per day). Relative liver weight was significantly increased in the 2000 mg/kg group only. Oxidative stress to the liver was shown by significant increase in SOD activity and the oxidative DNA-damage marker 8OHdG at all doses, as well as increased catalase (CAT) activity at 2000 mg/kg. However, no increase in DNA strand-breaks was observed by comet assay. Microarray analysis indicates that DP alters hepatic carbohydrate, lipid, nucleotide and energy metabolism as well as signal transduction processes.

105. In a 90-day oral study with rats exposed to DP at 0, 1, 10 or 100 mg/kg/d, DP preferentially accumulated in liver with *syn*-DP as the dominating stereoisomer (Li et al., 2013b). Clinical serum parameters such as alanine aminotransferase, aspartate aminotransferase and creatine kinase were reduced significantly at highest dose, and serum glucose significantly increased. A non-significant increase in thyroid stimulating hormone (TSH) were also observed. In the liver, gene-expression of several key enzymes were altered such as n-acetyltransferase, sulfotransferases and CYP2B1 (Li et al., 2013b). However, no significant changes in absolute body or liver weight or liver histopathology were observed.

106. Some of the earlier studies provided in Oxychem (2004b) indicate some liver and ovary effects. However, reliability of these studies is uncertain as discussed in ECHA (2017c) and description of the studies are also limited. In a 13-week oral feeding study in rats, absolute and relative liver weight was significant increased in males only, at 500 mg/kg, but no associated histopathology was observed (Oscarson, 1975, as described in Oxychem (2004b)). In a repeated-dose inhalation toxicity study, rats were exposed to 0, 640 or 1524 mg DP as dust/m³, 6 h per day, 5 days per week for 28 days (Bishop 1975). Significant increase in absolute and relative liver weight was observed at both doses. Corresponding hepatocytomegaly (swelling of liver cells with signs of cytotoxicity and necrosis) of centrolobular hepatocytes were observed for male rats at both doses and in some females at the higher dose. Effects on lung tissues were observed as increased number of macrophages in alveoli, and significant increase in absolute lung weight was observed for both doses in females and highest dose for males. A lowest-observed-adverse-effect concentration (LOAEC) of 640 mg/m³ was identified (Bishop 1975, as described in Oxychem 2004b). In a repeat-dermal study, male and female rabbits were dermally exposed to DP in 3% aqueous methylcellulose at 0, 500 or 2000 mg/kg bw/d, for 5 days per week, for 4 weeks on 20% of the total body surface (Trzyna 1975). None to minimal erythema occurred at the application site. In females, dose-related decrease in absolute and relative (to body and brain weight) ovary weight was observed at both doses, and a decrease in absolute and relative liver weight was observed at the highest dose. However, no histopathology effects were observed in any of the organs (Trzyna 1975, as described in OxyChem 2004b).

107. In a combined repeated-dose/reproductive/developmental toxicity screening test, conducted according to OECD guideline 422, CrI:CD (SD) rats were exposed to DP in corn oil at 0, 750, 1500 or 5000 mg/kg-bw/day by oral gavage (Brock et al. 2010). Animals (10/sex/dose) were treated for 28 days. No treatment-related effects were observed on clinical signs of toxicity, body weights, food consumption, neurobehavioral, and functional observational battery evaluations. No effects were observed on haematology, urinalysis, coagulation or clinical chemistry parameters. No dose-response related changes in organ weights (heart, liver, testes, ovaries, and thyroid/parathyroid glands) were observed. Mortalities and lesions were observed across all dose groups including controls, which were linked to gavage administration errors as indicated by gavage injury and administration of test substance in the thoracic/ pericardial cavity. The 5000 mg/kg vehicle in corn oil was described as particularly viscous. The authors identified a no-observable-effect level (NOEL) of 5000 mg/kg-bw/day. However, some effects was observed but considered not-treatment relevant by the authors since due to lack of dose-response relationship; A significant increase in absolute and relative ovary weight was observed in the 1500 mg/kg group only, and a significant increase in absolute and relative thyroid/parathyroid weight was observed in females in the 750 mg/kg group only. . A non-significant increase in number of mean early resorption sites (1.8) and postimplantation loss index (13.90%) in the 1500 mg DP/kg bw group compared to control (0.8 and 5.39%, respectively) were observed. Furthermore, pup body weight was significantly increased \geq 1500 mg/kg bw, mean pup weight was increased by 3, 15 and 13% compared to

control for the 750, 1500 and 5000 mg/kg groups. This effect may be a result as a result of DP's ability to activate the peroxisome proliferator-activated receptor PPAR- γ as described in paragraph below (Peshdary et al., 2019) as activation of this signal pathway has been linked to pregnancy/fetal outcomes by strengthening of uterus function (Hewitt et al., 2006; Díaz et al., 2012;), but also effects on the thyroid axis.

108. Adding to the above-mentioned indications of effects on thyroid hormones and liver metabolism, potential obesogenic effects have been observed for DP. In a mechanistic *in vitro* study, DP was shown to activate the peroxisome proliferator-activated receptor PPAR- γ and induce adipogenesis (observed as lipid accumulation and upregulation of adipogenesis mRNA and protein markers) of both murine and human preadipocytes. However, some DP-mediated adipogenic endpoints were independent of PPAR γ activation, suggesting that other potential modes of actions of DP may be involved (Peshdary et al., 2019).

109. Limited research has been performed towards the possible association between DP and the thyroid hormone (TH) homeostasis and lipoproteins in humans. An association between serum TH and DP levels was reported in human mother-infant pairs near an e-waste recycling area in China (Ben et al., 2014). DP concentration in maternal sera from the 20-year residents' group was 2 to 3-fold higher than the 3-year residents' group (geometric mean and range was 13.5, 1.28-900 and 3.68, 1.69-11.6 ng/g lw, respectively). *Syn*- and *anti*-DP were detected in placenta and umbilical cord serum samples, indicating that they could translocate from maternal to foetal tissues. Levels of TSH were significantly lower in maternal serum in the group that had lived in the area for 20 years (n= 44), than for those whose mothers had been residents to the area for 3 years or less (n=22), (p=0.046), but correlation analysis showed insignificant association between THs levels and DP levels in the 20-year residents' group. When concentrations of PBDEs were used as a control variable, the DP concentrations were positively associated with total triiodothyronine concentrations (TT3) in sera from mothers who had lived in the area for over 20 years (r = 0.37 and p = 0.020 for *syn*-DP, r = 0.360 and p = 0.024 for *anti*-DP). The association between serum DP levels and levels of the thyroid hormone TT3 suggested that DP may have some effects on thyroid hormone in humans (Ben et al., 2014). Another study compared serum levels of adults living in an e-waste recycling region (n=54) and a control region (n=58), lower levels of TSH, thyroid binding globulin and mRNA expression of thyroid receptor (TR α) and higher level of iodothyroine deiodinase 1 were observed in highly exposed individuals (Guo et al., 2019). The *syn*- and *anti*-DP mean and range in serum was 57 (12-1000) and 58 (11-1450) ng/g lw in residents from the e-waste recycling area, and 3.2 (0.36-12) and 5.9 (0.67-38) ng/g lw for the control group (Guo et al., 2019). Another study of pregnant women in Wenling, China observed no association between serum thyroid hormones and DP levels. The mean and range of DP in serum were 13.9 (0.41-252) ng/g lw (Yin et al., 2020). However, a positive relationship between DP serum level and apolipoprotein A was observed (p= 0.095) for total DP and 0.045 for *syn*-DP (Yin et al., 2020). A positive association (b= 0.089, p= 0.023) between *anti*-DP levels in the subjects' wrist band and blood levels of TSH was found for women (n= 62) in a study from the US (Wang et al., 2020).

Other concerns

110. The DP by-products 1,3- and 1,5-Dechlorane Plus monoadduct (DPMA) have in some cases been detected in greater levels than DP in environmental samples (Sverko et al., 2011; Tomy et al., 2013; Guerra et al., 2011), and also in Antarctic biota (Wolschke et al., 2015) (see paragraph 81 for details). DPMA may also be under-reported because of destructive sample preparation methods (Rjabova et al., 2016). Based on predictive QSAR modelling studies, DPMA is potentially both bioaccumulative and persistent. No information is available on its toxicity to humans and the environment, but based on its structural similarity to aldrin and heptachlor it might be epoxidated in the environment to form substances that could be neurotoxic and/or cause hepatotoxicity (ECHA 2017b,c). For additional information see Figure 1 and Table 14 in UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]

2. Synthesis of information

111. DP is a commercially available additive chlorinated FR that is marketed and used as a replacement for already regulated FRs in plastic polymers. Known applications include the aviation and automotive industries, electrical wire and cable coatings, plastic roofing materials, connectors in TV and computer monitors.

112. DP has been detected in dust, sludge and wastewater from WWTPs at sites far from production and e-waste recycling sites, indicating emission and exposure from consumer products. It is also detected in humans and wildlife. DP is a global contaminant and has been found in remote regions far from sites of production and use, including in the Arctic, the Antarctic and high-mountain regions in Tibet.

113. Available monitoring data provides evidence for ongoing LRET. Bioaccumulation and trophic magnification are reported in several studies, including in biota in remote regions. DP is environmentally persistent, and although existing environmental levels in remote regions are generally low, there is reason for concern. DP levels in remote regions are in some studies reported to be in the same range as in source regions. DP is persistent in the environment- and bioaccumulates in biota. The levels are likely to increase over time as a result of continued use and could possibly lead to adverse effects in biota in the future if no action is taken.

114. Although DP has been produced and used for almost half a century, the first environmental detection of DP was reported in 2006 in the Great Lakes Basin in North America (Hoh et al., 2006). Further research has identified DP as a global pollutant (Wang et al., 2016). Due to limited data, temporal and spatial trends for DP are equivocal; while increasing trends are observed in some studies, other studies report unchanged levels (see section 2.3.1).

115. Available short-term toxicity data indicates concern for potential adverse effects to the environment and humans. One relevant mode of action could be the induction of oxidative stress that influences several biological processes. Effects on the developing nervous system and brain have been indicated in fish. DP has been shown to activate peroxisome proliferator-activated receptor gamma, impair liver function and affect thyroid hormone pathway (see Table 5). Furthermore, a co-exposure study on zebrafish indicated that other pollutants may increase bioaccumulation, resulting in greater toxicity (Chen et al., 2019). Further data, including studies of longer duration and possibly different routes of exposure would provide better insight into the toxicity of DP. Structural similar analogue chemicals (mirex, chlordane, heptachlor, dieldrin/endrin, aldrin and endosulfane) are already listed under the Stockholm Convention (Table 1, UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]).

116. Adding to the concern for adverse effects are the DP by-products, such as 1,3- or 1,5-DPMA which have structural similarities to substances such as aldrin and heptachlor that are already listed by the Stockholm Convention (Table 14, UNEP/POPS/POPRC.16/INF/... [NOTE: INF IN PREPARATION]). There are no toxicity data for these monoadducts, but in some cases these are detected in greater levels than DP in environmental samples (Sverko et al., 2011; Tomy et al., 2013; Guerra et al., 2011).

117. DP is transferred to developing offspring during pregnancy via blood, and after delivery via breast feeding. Maternal transfer to eggs has been described for fish, bird and amphibians leading to exposure during sensitive life stages. High concentrations of DP have been detected in environmental samples and humans living near DP production plants and e-waste recycling sites.

118. Due to the POP properties and risks related to its widespread production and use, global action is warranted to prevent further releases of DP.

Table 5. POP characteristics of DP and its *syn*- and *anti*-isomer

Criterion	Meets the criterion (Yes/No)	Remarks
Persistence	Yes	<ul style="list-style-type: none"> Modelling of degradation potential and microbial metabolic pathways suggests that biodegradation is very slow and with low probability to degrade any faster than structural analogues that are listed under the Stockholm Convention (ECHA, 2017b; Zhang et al., 2016) Limited degradation in aging soil with only 4.2-8.2% of the initial DP having degraded after 260 days (Cheng et al., 2019) Limited degradation in soil over time (Wang et al., 2010a)
Bio-accumulation	Yes	<ul style="list-style-type: none"> Log K_{ow} of 9.3 (OxyChem, 2007) indicating high potential for bioaccumulation BCF >5000 (Wang et al., 2019) and depuration half-lives corresponding to a BCF >5000 (Tomy et al., 2008; Tang et al., 2018; Li et al., 2013b) BMFs/TMFs >1 reported for several species and food webs (Tomy et al., 2007; Wu et al., 2018; Sun et al., 2015, Sun et al., 2017; Kurt-Karakus et al., 2019; Na et al., 2017)
Potential for Long-Range Environmental Transport	Yes	<ul style="list-style-type: none"> DP is detected in the environment and biota in remote areas (Canada 2019a; AMAP 2017; Möller et al. 2010, 2012; Gao et al. 2018; Kim et al. 2018; Na et al. 2017; Yang et al. 2016; Liu et al. 2018). LRET occurs by DP sorption to particles in the atmosphere and in seawater (e.g. Möller et al., 2010; Möller et al., 2011; Möller et al., 2012; Sverko et al., 2011; Canada 2019a), and possibly also via migrating birds (Vorkamp et al. 2018). Concentrations in air decrease with increasing latitude (Möller et al. 2010, 2011).
Adverse effects	Yes	<ul style="list-style-type: none"> Indications of neurotoxicity in zebrafish (Chen et al., 2017; 2019) and carp (Li et al., 2019b; 2020). Potential for endocrine modulating effects in zebrafish (Kang et al., 2016), humans (Ben et al., 2014; Guo et al., 2019); and human-derived cells (Peshdary et al., 2019). Liver impairments as observed in mice and rats (Wu et al., 2012; Li et al., 2013b). Indication of immune modulating effects observed in carp (Li et al., 2019b).

Criterion	Meets the criterion (Yes/No)	Remarks
		<ul style="list-style-type: none"> • Oxidative stress in fish (Chen et al., 2017; Hang et al., 2013; Kang et al., 2016; Li et al., 2019b), marine bivalves (Barón et al., 2016, Gagne et al., 2017), earthworm (Zhang et al., 2014; Yang et al., 2016b), birds (Li et al., 2013a) and mice (Wu et al., 2012).

3. Concluding statement

119. DP is emitted into the environment from human activities, e.g. from manufacturing, use and disposal and management of waste. DP is persistent, bioaccumulative and undergoes LRET, making emissions and releases of this substance a transboundary pollution problem. Globally, DP is detected in humans, wildlife and environmental samples in all global regions, including in the Arctic and Antarctic.

120. In humans, DP has been detected in hair, umbilical cord blood, serum and breast milk. High concentrations of DP have been detected in environmental samples and humans living near e-waste recycling sites and production plants. DP is also detected in dust, sludge and wastewater from WWTPs indicating emission and exposure from consumer products throughout their lifecycle.

121. Available scientific literature suggests that there is a potential risk for adverse effects for developing aquatic species and organisms due to oxidative stress that affects important biological processes. The concern for adverse effects relates to observed effects on the liver, endocrine modulating effects, in particular to the thyroid hormone system and peroxisome proliferator-activated receptor gamma pathway. In aquatic organisms, effects on the developing nervesystem and brain have been indicated as well. Furthermore, observed effects on photosynthetic activity in marine macroalgae suggest possible impacts on aquatic ecosystems mediated by negative effects on primary production.

122. Mixture toxicity effects such as increased toxicity and bioaccumulation have been indicated for DP and suggest an additional cause for concern. Climate warming may exacerbate the mobilization and release of DP in polar environments as DP deposited in sea ice, glaciers, and permafrost are released into freshwater and marine environments. The combined effect of climate change and other environmental stressors adds to the risk by affecting exposure levels, the vulnerability and adaptability of organisms, particularly in the polar regions.

123. Based on evidence for persistence, bioaccumulation and adverse effects of DP observed in organisms and its widespread occurrence in the global environment including at remote regions, it is concluded that DP and its *syn*- and *anti*-isomers are likely, as a result of their long-range environmental transport, to lead to significant adverse human health and environmental effects such that global action is warranted.

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